Pro Glu Ala Pro Ala Vel Phe Ser Pro Glu Gly Thr Val Leu Leu Thr Gly Gly Thr Gly Ser Leu Gly Gly Leu Val Ala Lys Bis Leu Val Ala Arg His Gly Val Arg Arg Leu Val Leu Als Ser Arg Arg Gly Val Ala Ala Glu Asp Leu Val Thr Glu Leu Thr Glu Gin Gly Ala Thr Val Ser Val Val Ala Cys Asp Val Ser Asp Arg Asp Gin Val Ala Ala Leu Leu Als Glu His Arg Pro Thr Gly Ile Val His Leu Als Gly Leu Leu Asp Asp Gly Val lie Gly Ala Leu Asn Arg Glu Arg Leu Ala Gly Val Phe Ala Pro Lys Val Asp Ala Val Gln His Leu Asp Glu Leu Thr Arg Asp Leu Gly Leu Asp Ala Phe Val Val Phe Ser Ser Ala Ala Ala Leu Met Gly Ser Ala Gly Gln Gly Asn Tyr Ala Ala Asn Ala Phe Leu Asp Gly Let Met Ala Gly Arg Arg Ala Ala Gly Let Pro Gly Val Ser Let Als Trp Gly Leu Trp Glu Gln Ala Asp Gly Leu Thr Ala Asn Leu Ser

Als Thr Asp Gln Ala Arg Met Ser Arg Gly Gly Val Leu Pro Met Thr 1540 1545 1550

Pro Ala Giu Ala Leu Asp Ile Phe Asp Ile Gly Leu Ala Ala Glu Gln 1555 1560 1565

Ala Leu Leu Val Pro Ile Lys Leu Asp Leu Arg Thr Leu Arg Gly Gln 1570 1575 1580

Ala Thr Ala Gly Gly Glu Val Pro His Leu Leu Arg Gly Leu Val Arg 1585 1590 1595 1600

Als Ser Arg Arg Val Thr Arg Thr Als Als Als Ser Gly Gly Gly Gly 1605 1616

Leu Val His Lys Leu Ala Gly Arg Pro Ala Glu Glu Glu Glu Ala Val 1620 1625 1630

Lou Leu Gly Tle Val Gln Ala Glu Ala Ala Ala Val Leu Gly Phe Asn 1635 1640 1645

Ala Pro Glu Leu Ala Gin Gly Thr Arg Gly Phe Sar Asp Leu Gly Phe 1650 1655 1660

Asp Ser Leu Thr Ala Val Glu Leu Arg Asn Arg Leu Ser Ala Ala Thr 1665 1670 1675 1680

Gly Vai Lys Leu Pro Ala Thr Leu Val Phe Asp Tyr Pro Thr Pro Val 1685 1690 1693

Ala Leu Ala Arg His Leu Arg Glu Glu Leu Gly Glu Thr Val Ala Gly
1700 1705 1710

Als Pro Als Thr Pro Val Thr Thr Val Als Asp Als Gly Glu Pro Ile 1715 1720 1725

Ale Tie Val Gly Met Ale Cys Arg Leu Pro Gly Gly Val Met Ser Pro

AND AND Leu Trp Arg Met Val Ala Glu Gly Arg Asp Gly Met Ser Fro Phe Pro Cly Asp Arg Gly Trp Asp Leu Asp Gly Leu Phe Asp Ser Asp Pro Glu Ard Pro Gly Thr Ala Tyr Ile Arg Gln Gly Gly Phe Leu Ris Glu Ala Ala Leo Fhe Asp Pro Gly Phe Phe Gly Ile Ser Pro Arg Glu Ala Leu Ala Met Asp Pro Gln Gln Arg Leu Leu Ceu Glu Ala Ser Trp Glu Ala Leu Glu Arg Ala Gly Ile Asp Pro Thr Lys Ala Arg Gly Asp Ala Val Cly Val Phe Ser Cly Val Ser Ile His Asp Tyr Lew Clu Ser Leu Ser Asn Met Pro Ala Giu Leu Glu Gly Phe Val Thr Thr Ala Thr Ala Gly Ser Val Ala Ser Gly Arg Val Ser Tyr Thr Phe Gly Phe Glu Gly Pro Ala Val Thr Val Rap Thr Ala Cys Ser Ser Ser Leu Val Ala The His Leu Ala Ala Chn Ala Leu Arg Gln Gly Glu Cys Thr Mat Ala Let Ala Cly Gly Val Ala Val Mot Gly Ser Pro Ile Gly Val Ile Gly

Met Ser Arg Gln Arg Gly Met Ala Glu Asp Gly Arg Val Lys Ala Phe 1940 1945 1950

Ala Asp Gly Ala Asp Gly Thr Val Leu Ser Glu Gly Val Gly Ile Val 1955 1960 1965

Val Leu Glu Arg Leu Ser Val Ala Arg Glu Arg Gly His Arg Val Leu 1970 1975 1980

Ala Val Leu Arg Gly Ser Ala Val Asn Gln Asp Gly Ala Ser Asn Gly
1985 1990 2995 2000

Lew Thr Ala Pro Asn Gly Pro Ser Gln Gln Arg Val Ile Arg Ser Ala 2005 2010 2015

Lew Ala Gly Ala Gly Leu Gln Pro Ser Glu Vai Asp Val Val Glu Ala 2026 2025 2030

His Gly Thr Gly Thr Ala Leu Gly Glu Pro Ile Glu Ala Gln Ala Leu 2035 2040 2045

Leu Ala Thr Tyr Gly Lys Ser Arg Glu Thr Pro Leu Trp Leu Gly Ser 2050 2055 2060

Leu Lys Ser Asn Ile Gly His Thr Gln Ala Ala Ala Gly Val Ala Ala 2065 2070 2075 2080

Val Ile Lys Met Val Gin Ala Leu Arg Gin Asp Thr Leu Pro Pro Thr 2085 2090 2095

Leu Ris Val Gln Glu Pro Thr Lys Gln Val Asp Trp Ser Ala Gly Ala 2100 2105 2110

Val Glu Leu Thr Glu Gly Arg Glu Trp Ala Arg Asn Gly His Pro 2115 2120 2135

Arg Arg Ala Gly Val Ser Ser Phe Gly Ile Ser Gly Thr Asn Ala Ris 2136 2135 2140

Less lie Less Clu Clu Ala Pro Ala Asp Asp Thr Ala Glu Ala Asp Val 2145 2150 2155 2160

Pro Asp Ala Val Val Pro Val Val Ile Ser Ala Arg Ser Thr Gly Ser 2165 2170 2175

Leu Ala Gly Gin Ala Gly Arg Leu Ala Ala Phe Leu Asp Gly Asp Val 2180 2185 2190

Pro Leu Thr Arg Val Ala Gly Ala Leu Leu Ser Thr Arg Ala Thr Leu 2195 2200 2205

Thr Asp Arg Ala Val Val Ala Gly Ser Ala Glu Glu Ala Arg Ala 2210 2215 2220

Gly Leu Thr Ala Leu Ala Arg Gly Glu Ser Ala Ser Gly Leu Val Thr 2225 2230 2235 2240

Gly Thr Ala Gly Met Pro Gly Lys Thr Val Trp Val Phe Pro Gly Gln 2245 2250 2255

Gly Thr Gln Trp Ala Gly Met Gly Arg Glu Leu Leu Glu Ala Ser Pro 2260 2265 2270

Val Phe Ala Glu Arg Ile Glu Glu Cys Ala Ala Ala Leu Gln Pro Trp 2278 2280 2285

The Asp Trp Ser Lou Leu Asp Val Leu Arg Gly Glu Gly Glu Leu Asp 2290 2295 2300

Arg Val Asp Val Leu Gin Pro Als Cys Phe Als Val Met Val Gly Leu 2305 2310 2315 2320

Als Ala Val Trp Ala Ser Val Gly Val Val Pro Asp Ala Val Leu Gly

× 138 ×

His Ser Gln Gly Glu Ile Ala Ala Ala Cys Val Ser Gly Ala Leu Ser Leu Glu Asp Ala Ala Lys Val Val Ala Leu Arg Ser Gln Ala Ile Ala Ala Glu Leu Ser Gly Arg Gly Gly Met Ala Ser Ile Gln Leu Ser His Asp Glu Val Ala Ala Arg Leu Ala Pro Trp Ala Gly Arg Val Glu Ile Ala Ala Val Asn Gly Pro Ala Ser Val Val Ile Ala Gly Asp Ala Glu Als Leu Thr Glu Ale Val Glu Val Leu Gly Gly Arg Arg Val Ala Val Asp Tyr Ala Ser His Thr Arg His Val Glu Asp Ile Gln Asp Thr Lou Ala Glu Thr Leu Ala Gly Ile Asp Ala Gin Ala Pro Val Val Pro Phe Tyr Ser Thr Val Ala Gly Glo Trp Lie Thr Asp Ala Gly Val Val Asp Gly Gly Tyr Trp Tyr Arg Asn Leo Arg Asn Gln Val Gly Phe Gly Pro Ala Val Ale Glu Leu Ile Glu Gin Giy His Gly Vel Phe Val Glu Val Ser Ala His Pro Val Leu Val Gin Pro Ile Ser Glu Leu Thr Asp Ala

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Val Val Thr Gly Thm Leu Arg Arg Asp Asp Gly Gly Val Arg Arg Leu 2530 2535 2540

Less Thr Ser Met Ala Glu Less Phe Val Arg Gly Val Pro Val Asp Trp 2545 2550 2555 2560

Als Thr Met Ala Pro Pro Ala Arg Val Glu Leu Pro Thr Tyr Ala Phe 2565 2570 2575

Asp His Glo His Phe Top Leu Ser Pro Pro Ala Val Ala Asp Ala Pro 2580 2585 2590

Ala Leu Gly Leu Ala Gly Ala Asp His Pro Leu Leu Gly Ala Val Leu 2595 2600 2605

Pro Leu Pro Glin Ser Asp Gly Leu Val Phe Thr Ser Arg Leu Ser Val 2610 2615 2620

Arg Thr His Pro Trp Leu Ala Asp Gly Val Pro Ala Ala Ala Leu Val 2625 2630 2635 2640

Glu Leu Ala Val Arg Ala Gly Asp Glu Ala Gly Cys Pro Val Leu Ala 2645 2650 2655

Asp Leu Thr Val Glu Lys Leu Leu Val Leu Pro Glu Ser Gly Gly Leu 2665 2670

Arg Vel Cln Val Ile Val Ser Sly Glu Arg Thr Val Glu Val Tyr Ser 2675 2680 2685

Gln Let Glu Gly Alz Glu Asp Trp Ile Arg Asn Ala Thr Gly His Let 2690 2695 2700

Ser Ala Thr Ala Pro Ala Mis Glu Ala Phe Asp Phe Thr Ala Trp Pro 2705 2710 2715 2720

- Pro Ala Gly Ala Gln Gln Val Asp Gly Leu Trp Arg Arg Gly Asp Glu 2725 2730 2735
- Tie Phe Ala Giu Val Ala Leu Pro Giu Glu Leu Asp Aia Gly Ala Phe 2740 2745 2750
- Gly Ile His Pro Phe Leu Leu Asp Ala Ala Val Gln Pro Val Leu Ala 2755 2760 2765
- Asp Asp Glu Gla Pro Ala Glu Trp Arg Ser Leu Val Leu His Ala Ala 2770 2775 2780
- Gly Ala Ser Ala Leu Arg Val Arg Leu Val Pro Gly Gly Ala Leu Gln 2785 2790 2795 2800
- Ala Ala Asp Glu Thr Gly Gly Leu Val Leu Thr Ala Asp Ser Val Ala 2805 2810 2815
- Gly Arg Glu Leu Ser Ala Gly Lys Thr Arg Ala Gly Ser Leu Tyr Arg 2820 2825 2830
- Val Asp Trp Thr Glu Val Ser Ile Ala Asp Ser Ala Val Pro Ala Asn 2835 2840 2845
- Ile Glu Val Val Glu Ala Pne Gly Glu Glu Pro Leu Glu Leu Thr Gly 2850 2855 2860
- Arg Val Leu Glu Ala Val Gln Thr Trp Leu Val Thr Ala Ala Asp Asp 2865 2870 2875 2880
- Ala Arg Leu Val Val Thr Arg Gly Ala Val Arg Glu Val Thr Asp 2855 2890 2895
- Pro Ala Gly Ala Ala Val Trp Gly Leu Val Arg Ala Ala Gin Ala Glu 2900 2905 2910
- Asn Pro Gly Arg Ile Phe Leu Ile Asp Thr Asp Gly Glu Ile Pro Ala

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Leu Thr Gly Asp Glu Pro Glu Ile Ala Val Arg Gly Gly Lys Phe Phe Val Pro Arg Ile Thr Arg Ala Glu Pro Ser Gly Ala Ala Val Phe Arg Pro Asp Gly Thr Val Leu lie Ser Gly Ala Gly Ala Leu Gly Gly Leu Val Ala Arg Arg Leu Val Glu Arg His Gly Val Arg Lys Leu Val Leu Ala Ser Arg Arg Cly Arg Asp Ala Asp Gly Val Ala Asp Leu Val Ala App Leu Ala Ala Asp Val Ser Val Val Ala Cys Asp Val Ser Asp Arg Ala Gin val Ala Ala Leu Leu Asp Glu His Arg Pro Thr Ala Val Val His Thr Ala Gly Val Ile Asp Ala Gly Val Ile Glu Thr Leu Asp Arg Asp Arg Leu Ala Thr Val Phe Ala Pro Lys Val Asp Ala Val Arg His Leu Asp Giu Leu Thr Arg Asp Arg Asp Leu Asp Ala Phe Val Val Tyr Ser Ser Val Ser Ala Val Phe Met Gly Ala Gly Ser Gly Ser Tyr Ala Ala Ala Asn Ala Phe Leu Asp Gly Leu Met Ala Asn Arg Arg Ala Ala

Gly Leu Pro Gly Leu Ser Leu Ala Trp Gly Leu Trp Asp Gln Ser Thr 3125 3130 3135 Gly Met Ala Ala Gly Thr Asp Glu Ala Thr Arg Ala Arg Met Ser Arg 3140 3145 3150 Arg Gly Gly Leu Gln Ile Met Thr Gln Ala Glu Gly Met Asp Leu Phe 3155 3165 3160 Asp Ala Ala Leu Ser Ser Ala Glu Ser Leu Leu Val Pro Ala Lys Leu 3170 3175 3180 Asp Let Arg Gly Val Arg Ala Asp Ala Ala Ala Gly Gly Val Val Pro 3185 3190 3195 3200 His Met Leu Arg Gly Leu Val Arg Ala Gly Arg Ala Gln Ala Arg Ala 3205 3210 3215 Als Ser Thr Val Asp Asn Gly Leu Als Gly Arg Leu Ala Gly Leu Als 3220 3225 3230 Pro Ala Asp Gin Leu Thy Leu Leu Asp Leu Val Arg Ala Gin Val 3235 3240 3245

Ala Ala Val Leu Gly Sis Ala Asp Ala Ser Ala Val Arg Val Asp Thr 3250 3255 3260

Als Phe Lys Asp Als Gly Phe Asp Ser Leu Thr Als Val Glu Leu Arg 3265 3270 3275 3280

Asn Arg Met Arg Thr Ala Thr Gly Leu Lys Leu Pro Ala Thr Leu Val 3285 3290 3295

Phe Asp Tyr Pro Asn Pro Gln Ala Leu Ala Arg His Leu Arg Asp Glu 3300 3305 3310

Let Gly Gly Ala Ala Gin Thr Pro Val Thr Thr Ala Ala Ala Lys Ala 3325 3320 3325

Amp Leu Amp Glu Pro Ile Ala Ile Val Gly Met Ala Cym Arg Leu Bro 3330 3335 3340

Gly Gly Val Ala Gly Pro Glu Asp Leu Trp Arg Leu Val Ala Glu Gly
3345 3350 3355 3366

Arg Asp Ala Val Ser Ser Phe Pro Thr Asp Arg Gly Trp Asp Thr Asp 3365 3370 3375

Ser Lew Tyr Asp Pro Asp Pro Ala Arg Pro Gly Lys Thu Tyr Thu Arg 3380 3385 3390

His Gly Gly Phe Leu His Glu Ala Gly Leu Phe Asp Ala Gly Phe Phe 3395 3400 3405

Gly Ile Ser Pro Arg Glu Ala Val Ala Met Asp Pro Gln Gln Arg Leu 3410 3415 3420

Leu Leu Glu Ala Ser Trp Glu Ala Met Glu Asp Ala Gly Val Asp Pro 3425 3430 3435 3440

Let Ser Let Lys Gly Asn Asp Vel Gly Val Phe Thr Gly Met Phe Gly 3445 3450 3455

Gln Gly Tyr Val Ala Pro Gly Asp Ser Val Val Thr Pro Glu Leu Glu 3460 3465 3470

Gly Phe Ala Gly Thr Gly Gly Ser Ser Ser Val Ala Ser Gly Arg Val 3475 3480 3485

Ser Tyr Val Phe Gly Phe Glu Gly Pro Ala Val Thr Ile Asp Ser Ala 3490 3495 3500

Cys Ser Ser Ser Leu Val Ala Met His Leu Ala Ala Gln Ser Leu Arg

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Gin Gly Glu Cys Ser Met Ala Leu Ala Gly Gly Ala Thr Val Met Ala Asn Pro Gly Ala Phe Val Glu Phe Ser Arg Gln Arg Gly Lou Ala Val Asp Gly Arg Cys Lys Ala Phe Ala Ala Ala Ala Asp Gly Thr Gly Trp Ala Clu Gly Val Gly Val Val Ile Leu Clu Arg Leu Ser Val Ala Arg Glu Arg Gly His Arg Ile Leu Ala Val Leu Arg Gly Ser Ala Val Asn Gin Asp Gly Ala Ser Asn Gly Leu Thr Ala Pro Asn Gly Pro Ser Gln Gln Arg Val The Arg Arg Ala Leu Val Ser Ala Gly Leu Ala Pro Ser Asp Val Asp Val Val Glu Ala His Gly Thr Gly Thr Thr Leu Gly Asp Pro Ile Glu Ala Gln Ala Leu Leu Ala Thr Tyr Gly Lys Asp Arg Glu Ser Pro Leu Trp Leu Gly Ser Leu Lys Ser Asn Ile Gly His Ala Gln Ala Ala Ala Gly Val Ala Gly Val Ile Lys Met Val Gln Ala Leu Arg His Glu Val Leu Pro Pro Thr Leu His Val Asp Arg Pro Thr Pro Glu

Val Asp Trp Ser Ala Gly Ala Val Glu Leu Teu Teu Glu Ala Arg Glu 3715 3720 3725

Trp Pro Arg Asn Gly Arg Pro Arg Arg Ala Gly Val Ser Ala Phe Gly 3730 3740

Val Ser Gly Thr Asn Ala His Leu Ile Leu Glu Glu Ala Pro Ala Glu 3745 3750 3755 3760

Glu Pro Val Pro Thr Pro Glu Val Pro Leu Val Pro Val Val Val Ser 3765 3770 3775

Ala Arg Ser Arg Ala Ser Leu Ala Gly Gln Ala Gly Arg Leu Ala Gly 3780 3785 3790

Phe Val Ala Gly Asp Ala Ser Leu Ala Cly Val Ala Arg Ala Leu Val 3795 3800 3805

Thr Asn Arg Ala Ala Leu Thr Glu Arg Ala Val Met Val Val Gly Sex 3816 3815 3820

Arg Glu Glu Ala Val Thr Asn Leu Glu Ala Leu Ala Arg Gly Glu Asp 3825 3830 3835 3840

Pro Ale Ala Val Val Thr Gly Arg Ala Gly Ser Pro Gly Lys Leu Val 3845 3850 3855

Trp Val Phe Pro Gly Gln Gly Ser Gin Trp Ile Gly Met Gly Arg Glu 3860 3865 3870

Leu Leu Asp Ser Ser Pro Val Phe Ala Glu Arg Val Ala Glu Cys Ala 3875 3890 3885

Ala Ala Leu Glu Pro Trp Ile Asp Trp Ser Leu Leu Asp Val Leu Arg 3895 3900

Gly Glu Ser Asp Leu Leu Asp Arg Val Asp Val Val Gln Pro Ala Ser 3905 3910 3915 3920

Phe Ala Met Met Val Gly Leu Ala Ala Val Trp Gln Ser Val Gly Val
3925 3930 3935

Arg Pro Asp Ala Val Val Gly His Ser Gln Gly Glu Ile Ala Ala Ala 3940 3945 2950

Cys Val Ser Giy Ala Leu Ser Leu Gin Asp Ala Ala Lys Val Ala 3955 3960 3965

Leu Arg Ser Gln Ala Ile Ala Thr Arg Leu Ala Gly Arg Gly Gly Met
3970 3975 3980

Ala Ser Val Ala Leu Ser Glu Glu Asp Ala Thr Ala Trp Leu Ala Pro 3985 3990 3995 4000

Trp Ala Asp Arg Val Gin Val Ala Ala Val Asn Ser Pro Ala Ser Val 4005 4010 4015

Val Ile Ala Gly Glu Ala Gln Ala Leu Asp Glu Val Val Asp Ala Leu 4020 4025 4030

Ser Gly Gln Glu Val Arg Val Arg Val Ala Val Asp Tyr Gly Ser 4035 4040 4045

His Thr Asn Gln Val Glu Ala Ile Glu Asp Leu Leu Ala Glu Thr Leu 4050 4055 4060

Ala Gly Ile Glu Ala Gin Ala Pro Lys Val Pro Phe Tyr Ser Thr Leu 4055 4070 4075 4080

Ile Gly Asp Trp Ile Arg Asp Ale Gly Ile Val Asp Gly Gly Tyr Trp

4085 4090 4095

Tyr Arg Asn Leu Arg Asn Gin Val Gly Phe Gly Pro Ale Val Ala Glu

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Leu Val Arg Gln Gly His Gly Val Phe Val Glu Val Ser Ala His Pro Val Leu Val Gln Pro Leu Ser Glu Leu Ser Asp Asp Ala Val Val Thr Gly Ser Leu Arg Arg Glu Asp Gly Gly Leu Arg Arg Leu Leu Thr Ser Met Ala Glu Leu Tyr Val Glo Gly Val Pro Leu Asp Trp Thr Ala Val Let Pro Arg Thr Gly Arg Val Asp Let Pro Lys Tyr Als Phe Asp His Arg His Tyr Trp Leu Arg Pro Ala Glu Ser Ala Thr Asp Ala Ala Ser Leu Gly Gln Ala Ala Asp His Pro Leu Leu Gly Ala Val Val Glu Leu Pro Glm Ser Asp Gly Leu Val Phe Thr Ser Arg Leu Ser Val Arg The His Pro Try Leu Ala Asp His Ala Val Gly Gly Val Val Ile Leu Pro Gly Ser Gly Leu Ala Glu Leu Ala Val Arg Ala Gly Asp Glu Ala Gly Cys Thr Ala Leu Asp Glu Leu Ile Ile Glu Ala Pro Leu Val Val

Pro Ala Gin Gly Ala Val Arg Val Gin Val Ala Leu Ser Gly Pro Asp

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Glu Thr Gly Ser Arg Thr Val Asp Leu Tyr Ser Gln Arg Asp Gly Gly Als Gly Thr Trp Thr Arg His Ala Thr Gly Val Leu Ser Thr Ala Pro Ala Gin Giu Pro Glu Phe Asp Phe His Ala Trp Pro Pro Ala Asp Ala Glu Arg Ile Asp Val Glu Thr Phe Tyr Thr Asp Leu Ala Glu Arg Gly Tyr Gly Tyr Gly Pro Ala Phe Gln Gly Leu Gln Ala Val Trp Arg Arg Asp Gly Asp Val Phe Ala Glu Val Ala Leu Pro Glu Asp Leu Arg Lys Asp Ala Gly Arg Phe Gly Val His Pro Ala Leu Leu Asp Ala Ala Leu Gin Ala Ala Thr Ala Val Gly Gly Asp Glu Pro Gly Gln Pro Val Leu Ala Phe Ala Trp Asn Gly Leu Val Leu His Ala Ala Gly Ala Sor Ala Leu Arg Val Arg Leu Ala Pro Ser Gly Pro Asp Thr Leu Ser Val Ala Ala Ala Asp Glu Thr Gly Gly Leu Val Leu Thr Met Glu Ser Leu Val

Ser Arg Pro Val Ser Ala Glu Gln Leu Gly Ala Ala Ala Asp Ala Gly

His Asp Als Met Phe Arg Val Asp Trp Thr Glu Leu Pro Als Val Pro 4500 4505 4510

Arg Ala Glu Leu Pro Pro Trp Val Arg Ile Asp Thr Ala Asp Asp Val 4515 4520 4525

Ala Ala Leu Ala Glu Lys Ala Asp Ala Pro Pro Val Val Val Trp Glu 4530 4535 4540

Ala Ala Gly Gly Asp Pro Ala Leu Ala Val Ser Ser Arg Val Leu Glu 4545 4550 4555 4560

The Mat Gln Ala Trp Lou Ala Ala Pro Ala Phe Glu Glu Ala Arg Lou 4565 4570 4575

Val Val Thr Thr Arg Gly Ala Val Pro Ala Gly Gly Asp His Thr Leu 4580 4585 4590

Thr Asp Pro Ala Ala Ala Ala Val Trp Gly Leu Val Arg Ser Ala Gln 4595 4600 4605

Ala Glu His Pro Asp Arg Val Val Leu Leu Asp Thr Asp Gly Glu Val 4610 4615 4620

Pro Leu Cly Ale Val Leu Ale Ser Gly Glu Pro Gln Leu Ale Val Arg 4625 4630 4635 4640

Gly Thr Thr Phe Phe Val Pro Arg Leu Ala Arg Ala Thr Arg Leu Ser 4645 4650 4655

Asp Ala Pro Pro Ala Phe Asp Pro Asp Gly Thr Val Leu Val Ser Gly 4660 4670

Ala Gly Ser Leu Gly Thr Leu Val Ala Arg His Leu Val Thr Arg His 4675 4680 4685

Gly Val Arg Arg Val Val Leu Ala Ser Arg Gln Gly Arg Asp Ala Glu

Gly Ala Gln Asp Leu Tie Thr Glu Leu Thr Gly Glu Gly Ala Asp Val Ser Phe Val Ala Cys Asp Val Ser Asp Arg Asp Gin Val Ala Ala Leu Leu Ala Gly Leu Pro Asp Leu Thr Gly Val Val His Thr Ala Gly Val Phe Glu Asp Gly Val Ile Glu Ala Leu Thr Pro Asp Gln Leu Ala Asn \$755 Val Tyr Ale Ala Lys Val Thr Ala Ala Met His Leu Asp Glu Leu Thr Arg Asp Arg Asp Leu Gly Ala Phe Val Val Phe Ser Ser Val Ala Gly Val Met Gly Gly Gly Gla Gly Pro Tyr Ala Ala Ala Asn Ala Phe Leu Asp Ala Ala Met Ala Ser Arg Gin Ala Ala Gly Leu Pro Gly Leu Ser Leu Ala Trp Gly Leu Trp Glu Arg Ser Ser Gly Met Ala Ala His 484C Leu Ser Glu Val Asp His Ala Arg Ala Ser Arg Asn Gly Val Leu Glu Let The Arg Ala Glu Gly Let Ala Let Phe Asp Let Gly Let Arg Met

Ala Glu Ser Leu Leu Val Pro Ile Lys Leu Asp Leu Ala Ala Met Arg

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Ala Ser Thr Val Pro Val Leu Phe Arg Gly Leu Val Arg Pro Ser Arg
4900 4905 4910

Thr Gln Ala Arg Thr Ala Ser Thr Val Asp Arg Gly Leu Ala Gly Arg 4915 4920 4925

Len Ala Gly Len Pro Val Ala Glu Arg Ala Ala Val Len Val Asp Len 4930 4935 4940

Val Arg Gly Gln Val Ala Val Val Leu Gly Tyr Asp Gly Pro Glu Ala 4945 4950 4955 4960

Val Arg Pro Asp Thr Ala Phe Lys Asp Thr Gly Phe Asp Ser Leu Thr 4965 4970 4975

Ser Val Glu Leu Arg Asn Arg Leu Arg Glu Ala Thr Gly Leu Lys Leu 4980 4985 4990

Pro Ala Thr Leu Val Phe Asp Tyr Pro Asn Pro Leu Ala Val Ala Arg 4995 5000 5005

Tyr Lett Cly Ala Arg Lett Val Pro Asp Gly Thr Ala Asn Gly Asn Gly 5010 5015 5020

Asn Gly Asn Gly His Ser Glu Asp Asp Arg Leu Arg His Ala Leu Ala 5025 5030 5035 5040

Ale Ile Ale Ale Glu Asp Ale Gly Glu Glu Arg Ser Ile Ale Asp Leu 5045 5050 5055

Gly Val Asp Asp Leu Val Gln Leu Ala Phe Gly Asp Glu 5060 5065

- (2) IMPORMATION FOR SEQ ID NO: 6:
 - (i) SEQUENCE CHARACTERISTICS:

(A) LEWSTH: 1721 amino acids

(B) TYPE: amino acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:

Met Ala Cys Arg Leu Pro Gly Gly Val Thr Gly Pro Gly Asp Leu Trp
1 5 10 15

Arg Leu Val Ala Glu Gly Gly Asp Ala Val Ser Gly Phe Pro Thr Asp 20 25 30

Arg Cys Trp Asp Leu Asp Thr Leu Phe Asp Pro Asp Pro Asp His Ala 35 40 45

Gly Thr Ser Tyr Thr Asp Gln Gly Gly Phe Leu His Asp Ala Ala Leu 50 55 60

Phe Asp Pro Gly Phe Phe Gly Ile Ser Pro Arg Glu Ala Leu Ala Met 65 70 75 80

Asp Pro Gln Gln Arg Leu Leu Eeu Glu Ala Ser Trp Glu Ala Leu Glu 85 90 95

Gly Val Cly Leu Asp Pro Ala Ser Leu Gln Gly Thr Asp Val Gly Val
100 105 110

Phe Thr Gly Ala Gly Gly Ser Gly Tyr Gly Gly Gly Leu Thr Gly Pro 115 120 125

Glu Mot Gin Ser Phe Ala Gly Thr Gly Leu Ala Ser Ser Val Ala Ser

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	130					135					140				
Gly 145	Arg	Val	Ser	Tyr	Val 150	Phe	Gly	Phe	Glu	Gly 155	Pro	Ala	Val	Thr	Ile 160
Asp	Thr	Als	Cys	Ser 165	Ser	Ser	Læ	Val	Ala 170	Met	His	Leu	ala	Ala 175	Gln
Ala	Lec	Arg	Gln 180	Gly	Asp	Cys	Ser	Met 185	Ala	Leu	Ala	Gly	Gly 190	Ala	Met
Val	Met	Sex 195	Gly	210	Asp	Ser	Phe 200	Val	Val	Phe	Ser	Arg 205	Gln	Arg	Gly
Læ13	Ala 210	Thr	Asp	Gly	Yrg	Cys 215	Lys	Ala	Phe	Ala	Ser 220	Gly	Ala	Asp	Gly
Met 225	Val	Leu	Als	Glu	230 Gly	lie	Ser	Val	Val	val 235	Leu	Glu	Arg	Leu	Ser 240
Val	Ala	yrd	Glu	Arg 245	Gly	His	Arg	Val	Leu 250	alA	Val	Leu	Arg	Gly 255	Ser
ala	Val	Asn	Gln 260	Asp	Gly	ala	Ser	Asn 265	Gly	Læu	Thr	Ala	Pro 270	Asn	Gly
Bīo	Ser	Gln 275	Gln	Arg	Val.	lle	Arg 280	Ala	Ala	Leu	Ala	Asn 285	Ala	Gly	Ile
Gly	290 290	Sor	Assp	Val	Asp	Leu 295	Val	Glu	Ala	His	Gly 300	Thr	Gly	Thr	Ser
Leu 305	Gly	Ąsp	Pro	Tle	Glu 310	Ala	Gln	Ala	Len	Leu 315	Ale	Thr	#yr	Gly	Gln 320
Asp	Arg	Glu	The	Px0 325	Leu	Trp	Leu	Gly	Ser 330	Leu	Lys	Ser	Asn	Tle 335	Gly

His Thr Gln Ala Ala Ala Gly Val Ala Ser Val Ile Lys Val Val Gln

340

Ala Leu Arg His Gly Val Met Pro Pro Thr leu His Val Asp Glu Pro 355 360 365

345

350

Ser Ser Gln Val Asp Trp Ser Glu Gly Ala Val Glu Leu Thr Gly 370 380

Ser Arg Asp Trp Pro Arg Gly Asp Arg Pro Arg Arg Ala Gly Val Ser 385 390 395 400

Ser Phe Gly Val Ser Gly Thr Asn Val His Leu Ile Ile Glu Glu Ala 405 410 415

Pro Glu Glu Pro Ala Ala Ala Val Pro Thr Ser Ala Asp Val Val Pro 420 425 438

Leu Val Val Ser Ala Arg Ser Thr Gly Ser Leu Ala Gly Gln Ala Asp 435 440 445

Arg Leu Thr Glu Val Asp Val Pro Leu Gly His Leu Ala Gly Ala Leu 450 455 460

Val Ala Gly Arg Ala Val Lau Glu Glu Arg Ala Val Val Ala Gly 465 470 475 480

Ser Ala Glu Glu Ala Arg Ala Gly Leu Gly Ala Leu Ala Arg Gly Glu 485 490 495

Ala Ala Pro Gly Val Val Thr Gly Thr Ala Gly Lys Pro Gly Lys Val 500 505 510

Val Trp Val Phe Pro Gly Gln Gly Thr Gln Trp Val Gly Met Gly Arg 515 520 525

Glu	Leu 530	Leu	Asp	Ala	Ser	Pro 535	Val	Phe	Ala	Glu	Arg 540	Ile	Lys	Glu	Сух
Aì a 545	sîA	Ala	Leu	Asp	Gln 550	Trp	Thr	Asp	TTP	Ser 555	Leu	Leu	Asp	Val	Le u 560
Arg	Cly	Asp	Gly	Asp 565	Leu	Asp	Ser	Val	Glu 570	Val	Leu	Gln	Pro	Ala 575	Cys
Pho	Ala	Val	Met 580	Val	Gly	Leu	Ala	Ala 585	Val	Trp	Glu	Ser	Ala 590	Gly	Val
yrg	Pro	Asp 595	Ala	Val	Val	Gly	His 600	Ser	Gin	Gly	Glu	ïle 605	Ala	Ala	Ala
Суз	Val 610	Ser	Gly	Ala	Leu	The 615	ĭeu	Asp	Asp	Ala	Ala 620	Lys	Vel	val	Ala
Leu 625	Arg	Ser	Gln	Als	Ile 530	Ala	Ala	Arg	Leu	Ser 635	Gly	Arg	Gl.y	Gly	Met 640
Ala	Ser	Val	Ale	Leu 645	Ser	Glu	Asp	Gla	Ala 650	Asn	Ala	Arg	Lea	Gly 655	Leu
Try	Asp	Gly	Arg 660	Tle	Glu	Væl	Ala	Ala 665	Val	Asn	Gly	pro	Als 670	Ser	Val
Val	Tle	Ala 675	Gly	Asp	Ala	Gln	Ala 680	Leu	yad	Glu	Ala	Leu 685	Glu	Val	Leu
Ala	Gly 690	ąs¢	Gly	Val	Arg	Val 695	Arg	Gln	Val	ela	Val 700	Asp	Tyr	Ala	Ser
His 705	Thr	Arg	His	Val	Glu 710	Asp	Ile	Arg	Asp	Thr 715	Leu	Als	Glu	Thx	Leu 720

Ala Gly Ile Thr Ala Gln Ala Pro Asp Val Pro Phe Arg Ser Thr Val

915

				725					730					735	
Thr	Gly	Gly	Trp 740	Val	Arg	Asp	Ala	Asp 745	Val	Leu	Asp	Gly	Gly 750	Tyr	Trp
Tyr	Arg	Asn 755	Leu	Arg	Asn	Gln	Val 760	Arg	Phe	sly	Pro	Ala 765	Val	Ala	Gla
Læn	Leu 770	Glu	Gln	Gly	His	Gly 775	Val	Phe	Val	Glu	Val 780	Ser	Ala	His	Pro
Val 785	Lesta	Val	Gln	Pro	Tle 790	Ser	Glu	Leu	The	Asp 795	Als	Val	Val	Thr	800
Thr	Leu	Arg	Arg	Asp 805	Asp	Gly	Gly	Leu	Arg 810	Arg	Leu	Leu	Thr	Ser 815	Met.
Ale	Glu	Leu	Phe 820	Val	Arg	Gly	Val	Arg 825	Val	Äsp	Trp	Ala	Thr 830	Leti	Val
Pro	Pro	Ala 835	Arg	Val	Asp	Leu	Pro 840	Thr	Tyr	Ala	Phe	Asp 845	His	Gin	His
Phe	Trp 850	Leu	Arg	Pro	Ala	Ala 855	Gln	Ala	Asp	Als	Val 860	Ser	Leu	Gly	Gln
Ala 865	Ala	Ala	Glu	His	Pro 870	Leu	Leu	Gly	Ala	Val 875	Val	Arg	Leu	Pro	Gln 880
Ser	Asp	Gly	Leu	Val 885	Phe	Thr	šer	Ārģ	890 Tea	Ser	Leu	Ārģ	The	His 895	Pro
Try	Leu	alk	Asp 900	Bis	Thr	Ile	Gly	Gly 905	Val	Val	Len	Phe	Pro 910	Gly	Thr
Gly	Leu	Vel	Glu	Leu	Ale	Val	Arg	ala	Gly	Asp	Glu	Ala	Gly	Cys	Pro

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Val Leu Asp 930	Glu Leu	Val Thr 935	Glu Ala	Pro Leu	Val Val 940	Pro Gly	Gln
Gly Gly Val 945	Asn Val	Gln Val 950	Thr Val	Ser Gly 955	Pro Asp	Gln As n	Gly 960
Leu Arg Thr	Val Asp 965	Ile His	Ser Gln	Arg Asp 970	Asp Val	Trp Thr 975	Arg
His Ala Thr	Gly Thr 980	Val Ser	Ala Thr 985	Pro Ala	Ser Ser	Pro Gly 990	Phe
Asp Phe Thr 995	Ala Trp	Pro Pro	Pro Asp 1000	Gly Gln	Arg Val		Gly
Asp Phe Tyr 1010	Ala Asp	Leu Ala 101		Gly Tyr	Ala Tyr 1020	Gly Pro	Leu
Phe Gln Gly 1025	Val Arg	Ala Val 1030	Trp Gln	Arg Gly		Val Phe	Ala 1040
Glu Val Ala	Leu Pro 104		Arg Arg	Glu Asp 1050	Ala Ala	Arg Phe	
Leu His Pro	Ala Leu 1060	leu Asp	Ala Ala 106		Thr Gly	Thr Ile	Ala
Ala Ala Ala 107	***	Gln Pro	Gly Lys 1080	Ser Val	Met Pro 108		Trp

Asn Arg Leu Ala Leu His Ala Val Gly Ala Ala Gly Leu Arg Val Arg

Val Ala Pro Gly Gly Pro Asp Ala Leu Thr Val Giu Ala Ala Asp Glu

THE	Gly	Ala	Pro	Val	Leu	Pit	Met	qea	Ser	Leu	Ile	Leu	Arg	Glu	Val
				1125	Š				1130)				1135	ĝ

Als Leu Asp Gin Leu Asp Thr Als Arg Als Gly Ser Leu Tyr Arg Val 1140 1145 1150

Asp Trp Thr Pro Leu Pro Thr Val Asp Ser Ala Val Pro Ala Gly Arg 1155 1160 1165

Ala Glu Val Leu Glu Ala Phe Gly Glu Glu Pro Leu Asp Leu Thr Gly 1170 1175 1180

Arg Val Leu Ala Ala Leu Gln Ala Trp Leu Ser Asp Ala Ala Glu Glu 1185 1190 1195 1200

Alm Arg Leu Val Val Val Thr Arg Gly Ala Val Pro Ala Gly Asp Gly
1205 1210 1215

Val Val Ser Asp Pro Ala Gly Ala Ala Val Trp Gly Leu Val Arg Ala 1220 1225 1230

Ala Gln Ala Glu Asn Pro Asp Arg Phe Val Leu Leu Asp Thr Asp Gly 1235 1240 1245

Glu Val Pro Leu Glu Ala Val Leu Ala Thr Gly Glu Pro Gln Leu Ala 1250 1255 1260

Lew Arg Gly Thr Thr Phe Ser Val Pro Arg Lew Ala Arg Val Thr Glu 1265 1270 1275 1280

Pro Ala Glu Ala Pro Leu Thr Phe Arg Pro Asp Gly Thr Val Leu Val 1285 1290 1295

Ser Gly Ala Gly Thr Leu Gly Ala Leu Ala Ala Arg Asp Leu Val Thr 1300 1305 1310

Arg His Gly Val Arg Arg Len Val Leu Ala Ser Arg Arg Gly Arg Ala

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		1315	\$				1320	\$				1329	5		
Ala	Glu 1330		Il⊗	Asp	Asp	leu 1335		Ala	Glu	Leu	Thr 1340		His	Gly	Ala
Glu 134:	Vel :	Thr	Val	Ala	ala 1350		GRA.	Val	Ser	Asp 1355		Asp	Gln	Val	Ala 1360
Ala	Leu	Leu	Lys	Glu 1365		Ala	ĭeu	Thr	Ala 1370		Val	His	Thr	Ala 1375	
V&l	Phæ	Asp	Ala 1380		Val	The	Gly	Ala 1385		Thr	Arg	Glu	Arg 1390		είΑ
îys	Val	2he 1395		Pro	ľys	Val	Asp 140(Ala	Asn	His	Leu 1405		Glu	Lou
The Control	Arg 1410	,	Len	Ąsp	ī.eu)41:		Phe	Ile	Val	Tyr 1420		Ser	Ala	Ser
Ser 142	Ile i	The	Met.	Gly	Ale 1430		Ser	Gly	Gly	Tyr 1435		Ala	Ala	Asn	Ala 1440
Tyr	Iæu	Assp	Gly	Leu 1445		ala	Ala	Arg	Arg 1450		Ale	cly	leu	Pro 1455	
Let	Ser	Leu	Ais 1460		Gly	Pro	Trp	Glu 1465		Leu	Thr	Gly	Met 1470		Asp
The	Ile	Asp 1475		leu	Thi	Leu	Ala 1480		Net	Ser	Arg	Arg 148:		Gly	Arg
Gly	Gly 1490		Ārģ	Ala	Leu	Gly 149:		Ala	Asp	Gly	Met 1500		Leu	Phe	Asp
Ala 150:	Ala S	Leu	Als	Ala	Gly 1510		Ala	Leu	Leu	Val 1515		Ile	Glu	Leu	Asp 1520

Lou Arg Glu Val Arg Ala Asp Ala Ala Gly Gly Gly Thr Val Pro His 1525 1530 1535

Leu Leu Arg Gly Leu Val Arg Ala Gly Arg Gin Ala Ala Arg Thr Ala 1540 1545 1550

Ala Thr Glu Asp Gly Leu Glu Arg Arg Leu Ala Gly Leu Thr Val 1555 1560 1565

Ala Glu Glu Ala Leu Leu Leu Asp Leu Val Arg Gly Glu Val Ala 1570 1575 1580

Val Val Len Gly His Ala Asp Ser Ser Gly Val Arg Ala Asp Ala Ala 1585 1590 1595 1600

Phe Lys Asp Ala Gly Phe Asp Ser Leu Thr Ser Val Glu Leu Arg Asn 1605 1610 1615

Arg Leu Arg Glu Thr Thr Gly Leu Lys Leu Pro Als Thr Leu Val Phe 1620 1625 1630

Asp His Pro Asn Pro Leu Ala Leu Ala Arg His Leu Arg Ala Glu Leu 1635 1640 1645

Als Val Asp Glu Als Ser Pro Als Asp Als Val Leu Als Gly Leu Als 1650 1660

Cly Leu Glu Ala Ala Ile Ala Ala Ala Gly Ala Pro Asp Gly Asp Arg 1665 1670 1675 1680

Ile Thr Ala Arg Leu Arg Glu Leu Leu Lys Ala Ala Glu Aia Ala Glu
1685 1690 1695

Ala Arg Pro Gly Thr Ser Gly Asp Leu Asp Thr Ala Ser Asp Glu Glu 1700 1705 1710

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Law Phe Ala Leu Val Asp Gly Leu Asp 1715 1720

(2) INFORMATION FOR SEQ ID NO: 7:

(i) SEQUENCE CHARACTERISTICS:

(A) LEWSTH: 1688 amino acids

(B) TYPE: amino acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:

Met Ala Cys Arg Tyr Pro Gly Gly Val Ser Ser Pro Glu Asp Leu Trp 1 5 10 15

Arg Leu Val Ala Glu Gly Thr Asp Ala Val Ser Ala Phe Pro Gly Asp 20 25 30

Arg Gly Trp Asp Val Asp Gly Leo Val Asp Pro Asp Pro Asp Arg Pro 35 40 45

Gly Thr Thr Tyr Thr Asp Gln Gly Gly Phe Leu His Glu Ala Gly Leu 50 55 60

Phe Asp Ala Gly Phe Phe Gly Ile Ser Pro Arg Glu Ala Val Ala Met 55 70 75 80

Asp Pro Gin Gin Arg Leu Leu Giu Thr Ser Trp Glu Aia Ile Glu 85 90 95

Arg Thr Gly Thr Asp Pro Leu Ser Leu Lys Gly Ser Asp Ile Gly Val

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			100					105					110		
Phe	Thr	Gly 115	Val	Als	Ser	Met	Gly 120	Tyr	Gly	Ala	Gly	Gly 125	Gly	Val	Val
Als	Pro 130	Glu	Leu	Glu	Gly	Phe 135	Val	Gly	Thr	Gly	Ala 140	Als	Pro	Cys	ïle
Ala 145	Ser	Cly	Arg	Vai	Ser 150	Tyr	Val	Leu	Gly	Phe 155	Glu	Gly	Pro	Als	Val
Thr	Val	Asp	Thr	Gly 165	Cys	Ser	Ser	Ser	Leu 170	Val	Ala	Met	His	Leu 175	Ma
Ala	Gln	Ala	Leu 150	Arg	Arg	Gly	Els	Cys 185	Ser	Met	Ala	I&u	Ala 190	Gly	Gly
Ala	Met	Val 195	Met	Ala	Gln	Pro	200 Gly	Ser	Phe	Val	Ser	Phe 205	Ser	Arg	Gln
Ārģ	210	Leu	Ala	Leu	Asp	Gly 215	Arg	Cys	Lys	Ala	Phe 220	Set	Asp	Ser	Ala
Asp 225	Gly	Met	Gly	Leni	Ala 230	Gla	Gly	Val	Gly	Val 235	Ile	síA	Leu	Glu	Arg 240
Leu	Ser	Val	Aìa	Arg 245	Glu	Arg	Gly	His	Ar g 250	Val.	Leu	Ala	Val	Leu 255	Arg
Gly	Ile	Ala	Val 260	Asn	Gin	Asp	Gly	Ala 265	Ser	Asn	Cly	Leu	Thr 270	Alæ	Pro
Asn	Gly	Pro 275	Ser	Gln	Gln	Arg	Val 280	Ile	Arg	Ala	Ala	10u 205	Ala	Glu	Ala
Giy	Leu 290	Ser	Pro	Ser	Asp	Val 295	Asp	Ala	Val	Glu	Gly 300	His	Gly	Thr	Gly

Tax	Thr	Len	Gly	Asp	pro	Ile	Glu	Ala	Gln	Ala	i.eni	Lesu	Ala	The	Tyx
305					310					315					320
Gly	Lys	Gly	Arg	Asp 325	Pro	Glu	Lys	Pro	Iæu 330	TIP	Leu	Gly	Ser	Val 335	Lys
Ser	Asn	Leu	Gly 340	Ris	Zhr.	Gln	Als	Ala 345	Ala	Gly	Val	Ala	Ser 350	Val	Ile
Lys	Met	V&l 355	Gln	Als	Leu	Arg	His 360	Gly	Val	Leu	Pro	Pro 365	Thr	Leu	His
Val	As p 370	Arg	Pro	Sec	Thr	Glu 375	Val	Asp	Trp	Ser	Ala 380	Gly	Ala	Val	Sex
Leu 385	ren	The	Glu	Ala	Arg 390	Glu	Trp	Pro	årg	Glu 395	Gly	Arg	Pro	Arg	Arg 400
Ala	Gly	Væl	Ser	Ser 405	Phe	Gly	Il⊕	Ser	Gly 410	Thr	Asn	alA	His	Leu 415	Ile
Leu	Glu	Glu	Ala 420	Pro	Glu	Gžu	Glu	Pro 425	Pro	Väl	síñ	Glu	Ala 430	Pro	Ser
Ala	Gly	Val 435	Val	Pro	Val	Val.	Val 440	Ser	ala	Arg	Gly	Ala 445	Leu	Ala	Gly
Gln	Ala 450	Gly	Arg	Leu	Ala	Ala 455	Phe	leu	Glu	ala	Ser 460	Asz	Glu	Pro	ies:
Val 465	Thr	val	Als	Gly	Ala 470	Leu	lle	Cys	Gly	Azg 475	Ser	Arg	Phe	Gly	Asç 480
Arg	Ala	Val	Val	Val 485	Als	Gly	Thr	Arg	Ala 490	Glu	Als	Thr	Ala	Gly 495	I.eni

Ala	Ala	Leu	Ala 500	Arg	Gly	Glu	Ser	Ala 505	Ala	Asp	val	Val	Thr 510	Gly	Thi
Val	Ala	Ala 515	Ser	Gly	Val	Pro	Gly 520	Lys	Leu	Val	Trp	Val 525	Phe	Pro	Gly
Cln	Gly 530	Ser	Gln	TTP	Val	Gly 535	Met	Gly	Arg	Glu	Leu 540	Leu	Glu	aīa	Sex
Pro 545	Val	Phe	Ala	Ala	Arg 550	Ile	ala	Glu	Cys	Ala 555	Ala	sla	Leni	Glu	PTC 560
TYP	Ile	Asp	Trp	Ser 565	Leu	leu	Asp	Val	Leu 570	Arg	Gly	Glu	Gly	Asp 575	Let
Asp	Ai ÿ	Vzl	Asp 580	Val	Val	Gln	Pro	Ala 585	Ser	Phe	Ala	Val	Met 590	Val	Gly
Leu	Ala	Ala 595	Val	Trp	Ser	Ser	Val 600	Gly	Val	Val	Pro	Asp 605	Ala	Val	Leu
Gly	His 610	Ser	Gln	Gly	Glu	11e 615	ala	Ala	Ala	Cys	Val 620	Ser	Gly	sia	Le:
Ser 625	leu	Gla	Asp	Als	Ala 630	Lys	Val	Val	Ala	Leu 635	Arg	Ser	Gln	Ala	11e
Ala	Ala	Lys	1.eu	Ala 645	Gly	Arg	Gly	Gly	Met 650	Ala	Ser	Val	Ala	Leu 655	Ser
Glu	Glu	Asp	Als 660	Val	Ala	Arg	Leu	Arg 665	Kis	Trp	Ala	Asp	Arg 670	Val	Glu
Val	Ala	Ala 675	Val	Asn	Ser	Pro	Ser 680	Ser	Val	Val	Tìe	Als 695	Gly	Asp	Ala

Gir Ala Leu Asp Gin Ala Leu Giu Ala Leu Thr Gly Gin Asp Ile Arg

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	690					695					700				
Val 705	Arg	Arg	Val	Ala	Val 710	Asp	Tyr	Ala	ser	Mis 715	Thr	yrd	His	Val	Glu 720
Asp	lle	Gln	Glu	Pro 725	Leu	Ala	Glu	Ala	Len: 736	Ala	Gly	Ile	Glu	Ala 735	His
Ala	Pro	Thr	Leu 740	Pro	Phe	Phe	Ser	Thr 745	Leu	The	Gly	Asp	Trp 750	Ile	Arg
Glu	Ala	Gly 755	Val	Val	Asp	Gly	Gly 760	Tyr	Trp	Tyr	Arg	Asn 765	Leris	Arg	Asn
Gìn	Val 770	Gly	Phe	Gly	Pro	Ala 775	val	Ala	Glu	leu	leu 780	Gly	Leu	Gly	His
Arg 785	Val	Phe	Val	Glu	Val 790	Sex	Ala	His	Pro	Val 795	leu	Val	Gln	Ala	11e 800
Ser	Ala	Ile	Ala	Asp 805	Asp	The	Asp	Ala	V21 E10	Val.	Thr	Gly	Ser	Leu 815	ĀIĢ
Arg	Gla	Glu	Gly 820	Gly	Leu	Arg	Arg	Leu 825	leu	Mir	Ser	Met	Ala 830	Glu	Læu
Phe	Val	Arg 835	Gly	Val	Asp	Val	Asp 840	Trp	Ala	Thr	Met	Val 845	Pro	Pro	Ala
Arg	Val 850	Asp	Leni	Pro	Thr	Tyr 855	ÄÌä	Phe	Asp	His	Gln 860	His	Tyr	Trp	Leu
Arg 865	Tyr	Val	Glu	Thr	Ala 870	Thr	Ąsp	ala	Ala	Gly 875	Pro	Val	Val	Arg	Leu 880
Pro	Gln	Tha	Gly	Gly 885	Leu	Val	Fhe	Thr	Thr 890	Glu	Trp	Ser	Leu	Lys 895	Ser

Gln	Pro	Trp	Leu 900	Als	Glu	His	Thr	ieu 905	Glu	Asp	Leu	Val	Val 910	Val	Pro
Gly	Ala	Ala 915	Læu	Val	Glu	Leu	Ala 920	Val	Arg	Ala	Cly	As p 925	Glu	Ala	Gly
Thr	930	Val	Leu	qeA	Glu	Leu 935	Val	Ile	Glu	Thr	Pro 940	Leu	Val	Val	Pro
Glo 945	Arg	Gly	Ala	Ile	Arg 950	Val	Gln	Vel	Thr	Val 955	Ser	Gly	Pro	Asp	Asp 960
Gly	Thr	Arg	The	Leu 965	Glu	Val	His	Ser	Gln 970	Pro	Glu	Asp	Ala	Thr 975	Asp
Glu	TTP	Thr	ATG 980	His	Ala	Thr	G1y	Thr 985	Leu	Ser	Ala	The	Pro 990	Asp	Glu
Sør	Ser	Gly 995	Phe	Asp	Phe	The	Ala 1000		Pro	Pro	Pro	Gly 1005		Arg	Gln
Leti	As p 1010	-	Val	Pro	Ala	Ile 1011		Arg	Ala	Gly	Asp 1020		Ile	Phe	Ala
Glu 102:		Ser	Leu	Pro	asp 103(•	Ala	Asp	Ala	Glu 1035		Phe	Gly	Ile	Nis 1040
Pro	Als	Leu	Leu	Asp 104:	Ala 5	slA	Leu	His	Pro 1050		Leu	Pro	Gly	Asp 1055	_
Gly	Leu	Thr	Gîn 206(Met	Glu	Trp	Arg 1069	_	Leu	Thr	Leu	Eis 107(Ala
Gly	Ala	Ser	Thr	ĭ.eu	Arg	Val	Arg	rea	Val.	Pro	Gly	Gly	Phe	Læn	Glu

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1085

Ala Ala Asp Cly Ala Cly Ser Leu Val Val Thr Ala Lys Glu Val Ala 1090 1095 1100

Leu Arg Pro Val Thr Ile Ala Arg Ser Arg Thr Thr Thr Arg Asp Ser 1105 1115 1120

Leu Phe Gln Leu Asn Trp Tle Glu Leu Pro Glu Ser Gly Val Val Ala 1125 1130 1135

Ala Ala Asp Asp Thr Glu Val Leu Glu Val Pro Ala Gly Asp Ser Pro 1140 1145 1150

Leu Ala Ala Thr Ser Arg Val Leu Glu Arg Leu Gln Thr Trp Leu Thr 1155 1160 1165

Glu Pro Glu Ala Glu Gln Leu Vel Vai Vai Thr Arg Gly Ala Val Pro 1170 1175 1180

Ala Gly Asp Thr Pro Val Thr Asp Pro Ala Ala Ala Ala Val Trp Gly
1185 1190 1195 1200

Leu Val Arg Ser Ala Gin Ala Giu Asn Pro Asp Arg Ile Val Leu Leu 1205 1210 1215

Asp Thr Asp Gly Glo Val Pro Leu Gly Ala Val Leu Ala Gly Gly Glu 1220 1225 1230

Pro Gln Vai Ala Val Arg Gly Thr Ala Leu Tyr Val Pro Arg Leu Ala 1235 1240 1245

Arg Ala Asp Ala Ala Pro Val Ser Gly Leu His Gly Thr Val Leu Val 1250 1255 1260

Ser Gly Ala Gly Val Leu Gly Glu Ile Val Ala Arg His Leu Val Thr 1265 1270 1275 1280

Arg His Cly Val Arg Lys Leu Val Leu Ala Ser Arg Arg Cly Leu Asp

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Ala Asp Gly Ala Lys Asp Leu Val Thr Asp Leu Thr Gly Glu Gly Ala Asp Val Ser Val Val Ala Cys Asp Leu Ala Asp Arg Asn Gin Val Ala Ala Leu Leu Ala Asp His Arg Pro Ala Ser Val Ile His Thr Ala Gly Val Lou Asp Asp Gly Val Ile Gly Thr Lou Thr Pro Glu Arg Lou Ala Lys Val Phe Ala Pro Lys Val Asp Ala Val Arg His Leu Asp Glu Leu Thr Arg Asp Leu Asp Leu Asp Ala Phe Val Val Phe Ser Ser Gly Ser Gly Val Phe Gly Ser Pro Gly Gin Gly Asn Tyr Ala Ala Ala Asn Ala Phe Leu Aso Ala Ala Met Ala Ser Arg Arg Ala Ala Gly Leu Pro Gly Lou Ser Leu Ala Trp Gly Leu Trp Glu Gln Ala Thr Gly Met Thr Ala His Let Gly Gly Thr Asp Gln Ala Arg Met Ser Arg Gly Gly Val Arg Pro Ile Thr Ala Glu Glu Gly Met Ala Leu Phe Asp Thr Ala Leu Gly Ala Gin Pro Ala Leu Leu Val Pro Val Lys Leu Asp Leu Arg Giu Val

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Arg Ala Gly Gly Ala Val Pro Him Leu Leu Arg Gly Leu Val Arg Ala 1490 1495 1500

Giy Arg Arg Gin Ala Gin Ala Ala Ser Thr Val Asp Asn Gin Leu Leu 1505 1510 1515 1520

Gly Arg Leu Ala Gly Leu Gly Ala Pro Glu Gln Glu Ala Leu Leu Val 1525 1530 1535

Asp Leu Val Arg Gly Gin Val Ala Ala Val Leu Gly His Ala Gly Pro 1540 1545 1550

Asp Ala Val Arg Ala Asp Thr Ala Phe Lys Asp Ala Gly Phe Asp Ser 1555 1560 1565

Leu Thr Ser Val Asp Leu Arg Asn Arg Leu Arg Glu Ser Thr Gly Leu 1570 1575 1580

Lys Lett Pro Ala Thr Lett Ala Phe Asp Tyr Pro Thr Pro Lett Val Lett 1595 1590 1595 1600

Ala Arg His Leu Arg Asp Glu Leu Gly Ala Gly Asp Asp Ala Leu Ser 1605 1610 1615

Val Val His Ala Arg Leu Clu Asp Val Glu Als Leu Leu Gly Gly Leu 1620 1625 1630

Arg Leu Asp Glu Ser Thr Lys Thr Gly Leu Thr Leu Arg Leu Gln Gly 1635 1640 1645

Leu Val Ala Arg Cys Asn Gly Val Asn Asp Gin Thr Gly Gly Glu Thr 1650 1655 1660

Leu Ala Asp Arg Leu Glu Ala Ala Ser Ala Asp Glu Val Leu Asp Phe 1665 1670 1675 1680 - \$**70** -

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The Asp Glu Glu Leu Gly Leu Thr 1685

(2) INFORMATION FOR SEQ ID NO: 8:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 3413 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

Met Ala Thr Asp Glu Lys Leu Leu Lys Tyr Leu Lys Arg Val Thr Ala 1 5 10 15

Glu Leu His Ser Leu Arg Lys Glu Gly Ala Arg His Ale Asp Glu Pro 20 25 30

Le: Ala Val Val Gly Met Ala Cys Ary Phe Pro Gly Gly Val Sar Sar 35 40 45

Pro Glu Asp Leo Trp Gln Leo Val Ala Gly Gly Val Asp Ala Leo Ser 50 55 60

Asp Phe Pro Asp Asp Arg Gly Trp Glu Leu Asp Gly Leu Phe Asp Pro 65 70 75 80

Asp Pro Asp His Pro Gly Thr Ser Tyr Thr Ser Gln Gly Gly Phe Lew 85 90 95

Arg Gly Ala Gly Leu Phe Asp Ala Gly Leu Phe Gly Ile Ser Pro Arg

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Glu Ala Leu Val Met Asp Pro Gln Gln Arg Val Leu Leu Glu Thr Ser Trp Glu Ala Leu Glu Asp Ala Gly Val Asp Pro Leu Ser Leu Lys Gly Ser Asp Val Gly Val Phe Ser Gly Val Phe Thr Gln Gly Tyr Gly Ala Gly Ala The Thr Pro Asp Leu Glu Ala Phe Ala Gly Tie Gly Ala Ala Ser Ser Val Ala Ser Gly Arg Val Ser Tyr Val Phe Gly Leo Glo Gly Pro Ala Val Thr Ile Asp Thr Ala Cys Ser Ser Ser Leu Val Ala Ile His Leu Ala Ala Gln Ala Leu Arg Ala Gly Glu Cys Ser Met Ala Leu Ala Gly Gly Ala Thr Val Met Pro Thr Pro Gly Thr Phe Val Ala Phe Ser Arg Gin Arg Val Leu Ala Ala Asp Gly Arg Ser Lys Ala Phe Ser Ser Thr Ale Asp Gly Thr Gly Trp Ala Glu Gly Ala Gly Val Leu Val Leu Glu Arg Leu Ser Val Ala Gln Glu Arg Gly His Arg Ile Leu Ala Val Leu Arg Gly Ser Als Val Asn Gln Asp Gly Ala Ser Asn Gly Leu

Thr	Ala	Pro	Asn	Gly	Pro	ser	Gln	Gln	Arg	Val	ïïe	Arg	Lys	Alæ	Leu
305					310					315					320
Ala	Gly	äls	Gly	Leu	Val	Ala	Ser	qaA	Val	Ąsp	Val	Val	Glu	Ala	Ris
				325					330					335	
Gly	Thr	Gly	"tsr	Ala	Læsi	Gly	Asp	Pro	ĭle	Glu	Ala	Gln	Ala	Leu	Leu
			340					345					350		
Ala	Thr	Tvr	Glv	Gln	Glv	Arc	Glu	Arc	Pro	Læu	Tro	Lero	Glv	Ser	Va3
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žavs.	Ser	åæn	Pha	G) v	Nie	Mar.	Gle	Xla	\$la	a i a	a e	Va i	Δ\n	Gly	Wa I
w <u>r</u> w	370	X 300 X 3	30 31 81 60	-0404-35		375	10 di es	3:300.00	a acache	6 3.3.XE	380	E COLUM	*****	2023	****
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44 80 A					- 10 St. 10					48.57.48					400
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				405					410					415	
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			420					425					430		
							=								
Arg	Ala		Val	Ser	Ala	Phe		Ile	Ser	GTA	THE		Ala	Bis	Leu
		435					440					445			
lle	Leu	Glu	Glu	Ala	Pro	Pro	Ala	Asp	Ala	Val	ÄÌä	Glu	Glu	Pro	Glu
	450					455					460				
Phe	Lys	Gly	Pro	Val.	Pro	Ľėu	Val	Val	Ser	Ala	Gly	Ser	Pro	Tim	Ser
465					470					475					480
Leu	Ala	Ala	Gin	Ala	Gly	Arg	Len	Ala	Glu	Val	Leu	Ala	Ser	Gly	Gly
				485					490					495	

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Val	Ser	Axg	Ala	Arg	Leu	Als	Ser	Gly),en	Leu	Ser	Gly	Arg	Ala	Leu
			500					505					510		

- Leu Gly Asp Arg Ala Val Val Ala Gly Thr Asp Glu Asp Ala Val 515 520 525
- Ala Gly Leu Arg Ala Leu Ala Arg Gly Asp Arg Ala Pro Gly Val Leu 530 535 540
- The Gly See Ala Lys His Gly Lys Val Val Tyr Val Phe Pro Gly Gln 545 550 550 560
- Gly Ser Gln Arg Leu Gly Met Gly Arg Glu Leu Tyr Asp Arg Tyr Pro 565 570 575
- Val Phe Ala Thr Ala Phe Asp Glu Ala Cys Glu Gln Leu Asp Val Cys 580 585 590
- Leu Ala Gly Arg Ala Gly His Arg Val Arg Asp Val Val Leu Gly Glu 595 600 605
- Val Pro Ala Glu Thr Gly Leu Leu Asn Gln Thr Val Phe Thr Gln Ala 610 615 620
- Gly Leu Phe Ale Val Glu Ser Ale Leu Phe Arg Leu Ale Glu Ser Trp 625 630 635 640
- Gly Val Arg Pro Asp Val Val Leu Gly His Ser The Gly Glu The Thr 645 650 655
- Ala Ala Tyr Ala Ala Gly Val Phe Ser Leu Pro Asp Ala Ala Arg Ila 660 665 670
- Val Ala Arg Gly Arg Leu Met Gin Ala Leu Ala Pro Gly Gly Ala 675 680 685
- Met val Ala val Ala Ala Ser Glu Ala Glu val Ala Glu Leu Leu Gly

	690					695					700				
Asp 705	Gly	Val	Glu	Leu	Ala 710	Als	Val	Asn	Gly	Pro 715	Ser	Ala	Val	Val	1.ec
Ser	Gly	Asp	Ala	Asp 725	Ala	Val	Val	Als	Ala 730	Ala	Ala	Arg	Met	Arg 735	Glu
Arg	Gly	His	lys 740	Thr	Lys	Gln	leu	lys 745	Val	Ser	His	Als	Phe 750	His	Ser
Ala	Arg	Met 755	Ala	Pro	Met	Leu	Ala 760	Glu	Phe	Ala	Ala	Glu 765	Leu	Ala	Gly
Val	Thr 770	Tr	yrd	Glu	Pro	Glu 775	Ile	Pro	Val	Val	Ser 780	Asn	Val	The	Gly
Arg 785	Phe	Ala	Glu	Pro	Gly 790	Glu	Leu	Thr	Glu	Pro 795	Gly	Tyr	Trp	Als	Glu 800
His	Val	Arg	Ang	805 Pro	Val	Arg	Phe	Ala	Glu 810	Gly	Val	Ala	Ala	Als 815	Thr
Glu	Ser	Gly	Gly 820	Ser	Leu	Phe	Val	Glu 825	Leu	Gly	Pro	Gly	Ala 008	Ala	Leu
Thr	alA	Leu 835	Val	Glu	Glu	Thr	Ala 840	Glu	Val	Thr	Cys	Val 845	Ala	Ala	Len
şrg	Asp 850	Asp	Arg	Pro	Glu	Val 855	Thr	Ala	Leu	Ile	Thr 860	Ala	Val	Ala	Glu
Leu 965	Phe	Val	Arg	Gly	Val 870	Ala	Val	Ąsp	zib	Pro 875	Ala	Leu	læu	Pro	Pr0 880
Val	**************************************	Gly	Fhe	Val 885	Asp	Leu	Pro	Lys	Tyr 890	Ala	Phe	Asp	Gln	Gln 895	His

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Tyr Trp Leu Gln Pro Ala Ala Gin Ala Thr Asp Ala Ala Ser Leu Gly
900 905 910

Gln Val Ala Ala Asp His Pro Leu Cly Ala Val Val Arg Leu Pro 915 920 925

Gln Ser Asp Gly Leu Val Phe Thr Ser Arg Leu Ser Leu Lys Ser His 930 935 940

Pro Trp Leu Ala Asp His Val Lie Gly Gly Val Val Leu Val Ala Gly 945 955 960

Thr Gly Leu Val Glu Leu Ala Val Arg Ala Gly Asp Glu Ala Gly Cys 965 970 975

Pro Val Leu Glu Glu Leu Val Ile Glu Ala Pro Leu Val Val Pro Asp 980 985 990

His Gly Gly Val Arg Ile Gln Val Val Val Gly Ale Pro Gly Glu Thr 595 1000 1005

Gly Ser Arg Ala Val Glu Vel Tyr Ser Leu Arg Glu Asp Ala Gly Ala 1010 1015 1020

Glu Val Trp Ala Arg His Als Thr Cly Pha Lau Ala Ala Thr Pro Ser 1025 1030 1035 1040

Gin His Lys Pro Phe Asp Phe Thr Ala Trp Pro Pro Pro Gly Vel Glu 1045 1050 1055

Arg Val Asp Val Glu Asp Phe Tyr Asp Gly Leu Val Asp Arg Gly Tyr 1060 1065 1070

Ala Tyr Gly Pro Ser Phe Arg Gly Leu Arg Ala Val Trp Arg Arg Gly 1075 1080 1085 WO 98/07868

qa4	Glu 1090		Phe	Ala	Glu	Val	Ala j	Leu	Ala	Glu	As p	-	Arg	Ala	Asp
Ala 1105		Arg	Phe	G1y	Ile 1116		Pro	Gly	Leu	Leo 111:		Ala	Ala	Leu	His 1120
Ala	Gly	Met	Ala	Gly 1125		Thr	Thr	Thr	Glu 1130		Pro	Gly	Arg	Pro 113f	
Leni	Pro	Phe	Ala 1140		Asn	Gly	Leu	Val 1145		Bis	Ala	Ala	Gly 1150		Ser
Ăla	Leu	Arg 1155		Arg	Leu	Ala	Pro 1160		Gly	Pro	Asp	Ala 1165		ser	Val
Glu	Al a 1170		Asp	Glu	Ala	Gly 1175	Gly ;	Leu	vsl	Val	Thr 1180		qzA	Ser	Leu
Val 1185		Arg	Pro	Val	5er 1190		Glu	Gln	Leu	Gly 1195		Als	Åla	Äsn	His 1200
Asp	Ala	Leu	Phe	Arg 1205		Glu	Trp	reir	Glu 1210		Ser	Ser	Ala	Gly 1215	
Val	Pro	Ale	Asp 1220		Val	Glu	Val	Leu 1225		Ala	Val	Gly	Glu 1230		Pro
læn	Glu	Leu 1235		Gly	Arg	Val	Leu 1240		Ala	Val	Gln	Thr 1245		Leu	Aĩa
Asp	Ala 1250		Asp	ązá	Als	Ar ç 1255	Leu	Val	Val	Val	Thr 1260	••	Gly	Ala	Väl
His 1265		Væl	Thr	Asp	Pro 1270		Gly	Ala	Ala	Val 1275		Gly	Leu	Ile	Arg 1280

Ala Ala Gln Ala Glu Asn Pro Asp Arg Ile Val Leu Leu Asp Thr Asp

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Gly Glu Val Pro Leu Gly Arg Val Leu Ala Thr Gly Glu Pro Gln Thr Ala Val Arg Gly Ala Thr Leo Phe Ala Pro Arg Leo Ala Arg Ala Glo Ala Ala Glu Ala Pro Ala Val Thr Gly Gly Thr Val Lau Ile Ser Gly Ala Gly Ser Leu Gly Ala Leu Thr Ala Arg His Leu Val Ala Arg His Gly Val Arg Arg Lew Val Lew Val Ser Arg Arg Gly Pro Asp Ala Asp Gly Met Ala Glu Leu Thr Ala Glu Leu Ile Ala Gln Gly Ala Glu Val Ala Val Val Ala Cys Asp Leu Ala Asp Arg Asp Gln Val Arg Val Leu Leu Ala Clu His Arg Pro Asn Ala Val Val His Thr Ala Gly Val Leu Asp Asp Gly Val Phe Glu Ser Lou Thr Arg Glu Arg Lou Ala Lya Val Phe Ala Pro Lys Val Thr Ala Ala Asn Bis Leu Asp Glu Leu Thr Arg Glu Leu Asp Leu Arg Ala Phe Val Val Phe Ser Ser Ala Ser Gly Val

Phe Gly Ser Ala Gly Gin Gly Asn Tyr Ala Ala Ala Asn Ala Tyr Leu

Asp	Ala	Val	Val	Ala	Asn	Arg	Arg	Ala	Ala	Gly	Leu	Pro	Gly	Thr	Ser
	149)				149	Ş				150	0			
Leu	Ala	TTP	Gly	Leu	TTP	Glu	Gln	Thr	Asp	Gly	Met	The	Ala	His	Leu
1509	ŝ				1510)				1513	5				1520
Gly	Asp	Als	Asp	Gln	Ala	Arg	Ala	Ser	Arg	Gly	Gly	Val	Leu	Ala	lle
				1525	Š				1530)				1533	Ĵ
2er	Pro	Ala	Glu	Gly	Met	Glu	Leu	Phe	qea	Ala	Ala	Pro	Asp	Gly	Leu
			1540)				1543	\$				1550)	
Val	Val	pro	Val	Lys	Leu	Asp	Leu	Arg	Lys	Thr	Arg	Ala	Gly	Gly	Thr
		1555	<u> </u>				1560)				1563	3	_	
Val	Pro	His	Lesi	Leu	Arg	Gly	Leu	Val	Arg	Pro	Gly	Arg	Gln	Gln	Ala
	1570)				1575	Š				1580)			
Arg	Pro	Ala	Ser	Thr	Važ	Asp	Asn	Gly	Len	Ala	Gly	Arg	Leu	Ala	Gly
1585					1590					1595	••				1600
Leu	Ala	Pro	Ala	Glu	Gln	Glu	Ala	Leu	Leu	Len	Asp	Vai	Val	Arg	Thr
				1605	à				1610	}	_			1615	į
Gln	Val	Ala	Leu	Val	[æii	Gly	His	Ala	Gly	Pro	Glu	Ala	Val	Aro	Alæ
			1620			-30		1625	-				1630	-	
caA	Thr	sí£	Phe	Ive	Aso	Thr	Glv	Phe	Aso	Ser	Leu	Thr	Sec	Val	Glu
-		1633					1640		¥.			1645			
Leu	Ang	Asn	Arg	Lea	Arc	Glu	Ala	Ser	Glv	Leu	Lvs	Ĩæu	Pro	Ala	Thr
	1650		#*		*	1655			- 4		1660				
	•										-4E-15-17				
Leu	Val	Phe	Asp	Tyr	org	The	Pro	Val	Ala	Leu	Ala	Arq	TVI	Leu	Airc
													177		

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Asp Glu Leu Gly Asp Thr Val Ala Thr Thr Pro Val Ala Thr Ala Ala 1685 1690 1695

Ala Ala Asp Ala Gly Glu Pro Ile Ala Ile Val Gly Met Ala Cys Arg 1700 1705 1710

Lem Pro Gly Gly Val Thr Asp Pro Glu Gly Leu Trp Arg Leu Val Arg 1715 1720 1725

Asp Gly Leu Glu Gly Leu Ser Pro Phe Pro Glu Asp Arg Gly Trp Asp 1730 1735 1740

Less Glu Asn Les Phe Asp Asp Asp Pro Asp Arg Set Gly Thr Thr Tyr 1745 1750 1755 1760

Thr Ser Arg Gly Gly Phe Leu Asp Gly Ala Gly Leu Phe Asp Ala Gly 1765 1770 1775

Phe Phe Cly Ile Ser Pro Arg Glu Ala Leu Ala Met Asp Pro Gln Gln 1780 1785 1790

Arg Leu Leu Glu Ala Ala Trp Glu Ala Leu Glu Gly Thr Gly Val 1795 1800 1805

Asp Pro Gly Ser Lew Lys Gly Ala Asp Val Gly Val Phe Ala Gly Val 1810 1815 1820

Ser Asn Gln Cly Tyr Gly Met Gly Als Asp Pro Ala Giu Leu Ala Giy 1825 1830 1835 1840

Tyr Ala Ser Thr Ala Gly Ala Ser Ser Val Val Ser Gly Arg Val Ser 1845 1850 1855

Tyr Val Phe Gly Phe Glu Gly Pro Ala Val Thr Ile Asp Thr Ala Cys 1860 1865 1870

Ser Ser Ser Leo Val Ala Met Him Leo Ala Gly Gin Ala Leo Arg Gin

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		1873	Š				198)				188	5		
Gly	Glu 189(Ser	Met	Ala	L@C		Gly	gly	Val	Thr 190		Met	Gly	The
Pro 1905		Thr	Phe	Val	Glu 191(Ala	Lys	Gln	Arg 191:		Len	Ala	Gly	As p 1920
Gly	Arg	Cys	Lys	Ala 1925		Ala	Glu	Gly	Ala 193(-	Gly	Thr	Gly	Trp 1935	
Glu	Gly	Val	Gly 1940		Val	Val	Leu	Glu 1945		Len	Ser	Val	Ala 1950		Glu
ĀIĢ	Gly	His 1959	Arg i	Val	Leu	Ala	Val 1960		Arg	Gly	Ser	Ala 1969		Asn	Ser
Asp	Gly 1970		Ser	Asn	Gly	Leu 1975		Ala	Pro	Asn	Gly 198(Ser	Gln	Gln
Arg 1985		Tle	Arg	Arg	Ala 1990		Aìa	Gly	Ala	Gly 1995		Glu	Pro	Ser	As p 2000
Val	Asp	lie	Val	Glu 2005	**	His	Gly	Thr	Gly 2010		Ala	Leu	Gly	Asp 2015	
ïle	Glu	Ala	Gln 2020		Leu	Leu	Ala	Thr 2025		Gly	Lys	Asp	Arg 2030		Pro
Glu	Thr	Pro 2035	Leu ;	TTP	Leu	Gly	Ser 204(Lys	Ser	Asn	Phe 2045	•	His	Thr
Gln	Ser 2050		Ala	Gly	Val	Ala 2055	•	Val	lle	Lys	Met 2060		Gln	Ala	Leu
Arg	His	Gly	Va.l	Met	Pro	pro	Thr	Leu	His	val	qeA	Arg	Pro	Thr	Ser

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Gin Val Asp Trp Ser Ala Gly Ala Val Glu Val Leu Thr Glu Ala Arg 2085 2090 2095

Glu Trp Pro Arg Asn Gly Arg Pro Arg Arg Ala Gly Val Ser Ser Phe 2100 2105 2110

Gly The Ser Gly Thr Asn Ala His Leu Ile Ile Glu Glu Ala Pro Ala 2115 2120 2125

Glu Pro Gln Leu Ala Gly Pro Pro Pro Asp Gly Gly Val Val Pro Leu 2130 2135 2140

Val Val Ser Ala Arg Ser Pro Gly Ala Lau Ala Gly Gln Ala Arg Arg 2145 2150 2155 2166

Leu Ala Thr Phe Leu Gly Asp Gly Pro Leu Ser Asp Val Ala Gly Ala 2165 2170 2175

Leu Thr Ser Arg Ala Leu Phe Gly Glu Arg Ala Val Val Val Ala Asp 2180 2185 2190

Ser Ala Glu Glu Ala Arg Ala Gly Leu Gly Ala Leu Ala Arg Gly Glu 2195 2200 2205

Asp Ala Pro Gly Leo Val Arg Cly Arg Val Pro Ala Ser Gly Leo Pro 2210 2215 2220

Gly Lys Leu Val Trp Val Phe Pro Gly Gln Gly Thr Gln Trp Val Gly 2225 2230 2235 2240

Met Gly Arg Glu Leu Leu Glu Glu Ser Pro Val Phe Ala Glu Arg Ile 2245 2250 2255

Ala Glu Cys Ala Ala Ala Leu Glu Pro Trp Ile Gly Trp Ser Leu Phe 2260 2265 2270

- Asp Val Leu Arg Gly Asp Gly Asp Leu Asp Arg Val Asp Val Leu Gln 2275 2280 2285
- Pro Ala Cys Phe Ala Val Met Val Gly Leu Ala Ala Val Trp Ser Ser 2290 2295 2300
- Ala Gly Val Val Pro Asp Ala Val Leu Gly His Ser Gln Gly Glu Ile 2305 2310 2315 2320
- Ala Ala Cys Val Ser Sly Ala Leu Ser Leu Glu Asp Ala Ala Lys 2325 2330 2335
- Val Val Ala Leu Arg Ser Glm Ala Ile Ala Ala Lys Leu Ser Gly Arg 2340 2345 2350
- Gly Gly Met Ala Ser Val Ala Leu Gly Glu Ala Asp Val Val Ser Arg 2355 2360 2365
- Leu Ala Asp Gly Vai Glu Vai Ala Ala Val Asn Gly Pro Ala Ser Vai 2370 2375 2380
- Val Tie Ala Gly Asp Ala Gln Ala Leu Asp Glu Thr Leu Glu Ala Leu 2385 2390 2395 2400
- Ser Gly Ala Gly Ila Arg Ala Arg Arg Val Ala Val Asp Tyr Ala Ser 2405 2410 2415
- His Thr Arg His Val Glu Asp Ile Glu Asp Thr Leu Ala Glu Ala Leu 2420 2425 2430
- Ala Gly Ile Asp Ala Arg Ala Pro Leu Val Pro Phe Leu Ser Thr Leu 2435 2440 2445
- The Gly Glu Trp Ile Arg Asp Glu Gly Val Val Asp Gly Gly Tyr Trp 2450 2455 2460
- Tyr Arg Asn Leu Arg Gly Arg Val Arg Phe Gly Pro Ala Val Glu Ala

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Leu Leu Ala Gin Gly His Gly Val Phe Val Giu Leu Ser Ala His Pro Val Leu Val Glm Pro Ile Thr Glu Leu Thr Asp Glu Thr Ala Ala Val Val Thr Gly Ser Leu Arg Arg Asp Asp Gly Gly Leu Arg Arg Leu Leu Thr Ser Mer Ala Glu Leu Phe Val Arg Gly Val Glu Val Asp Trp Thr Ser Les Val Pro Pro Ala Arg Ala Asp Les Pro Thr Tyr Ala Phe Asp His Glu His Tyr Trp Leu Arg Ala Ala Asp Thr Ala Ser Asp Ala Val Ser Leu Sly Leu Ala Gly Ala Asp His Pro Leu Leu Gly Ala Val Val Gin Leu Pro Gin Ser Asp Gly Leu Val Phe Thr Ser Arg Leu Ser Leu Arg Ser His Pro Trp Leu Als Asp His Ala Vel Arg Asp Val Val Ile Val Pro Gly Thr Gly Leu Val Glu Leu Ala Val Arg Ala Gly Asp Glu Ala Gly Cys Pro Val Leu Asp Glu Leu Val Ile Glu Ala Pro Leu Val

Val Pro Arg Arg Gly Gly Val Arg Val Gln Val Ala Leu Gly Gly Pro

- Als Asp Asp Gly Ser Arg Thr Val Asp Val Phe Ser Leu Arg Glu Asp 2675 2680 2685
- Als Asp Ser Trp Leu Arg His Als Thr Gly Vel Leu Val Pro Glu Asn 2690 2695 2700
- Arg Pro Arg Gly Thr Als Als Phe Asp Phe Als Als Trp Pro Pro Pro 2705 2716 2715 2720
- Glu Ala Lys Pro Val Asp Leu Thr Gly Ala Tyr Asp Val Leu Ala Asp 2725 2730 2735
- Val Gly Tyr Gly Pro Thr Phe Arg Ala Val Arg Ala Val Trp 2740 2745 2750
- Arg Arg Gly Ser Gly Asn Thr Thr Glu Thr Phe Ala Glu Ile Ala Leu 2755 2760 2765
- Pro Glu Asp Ala Arg Ala Glu Ala Gly Arg Phe Gly Ile Nis Pro Ala 2770 2775 2780
- Leu Leu Asp Ala Ala Leu Nis Ser Thr Met Val Ser Ala Ala Ala Asp 2785 2790 2795 2800
- The Glu Ser Tyr Gly Asp Glu Val Arg Leu Pro Phe Ala Trp Asn Gly 2805 2810 2815
- Leu Arg Leu His Ala Ala Gly Ala Ser Val Leu Arg Val Arg Val Ala 2820 2825 2830
- Lys Pro Glu Arg Asp Ser Leu Ser Leu Glu Ala Val Asp Glu Ser Gly 2835 2840 2845
- Gly Leu Val Val The Leu Asp Ser Leu Val Gly Arg Fro Val Ser Asn 2850 2855 2860

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Asp Gln Leu Thr Thr Ala Ala Gly Pro Ala Gly Ala Gly Ser Leu Tyr 2865 2870 2875 2880

Arg Val Asp Trp Thr Pro Leu Ser Ser Val Asp Thr Ser Gly Arg Val 2885 2890 2895

Pro Ser Trp Leu Pro Val Ala Thr Ala Glu Glu Val Ala Thr Leu Ala 2900 2905 2910

Asp Asp Val Leu Thr Gly Ala Thr Glu Ala Pro Ala Val Ala Val Met 2925 2920 2925

Glu Ala Val Ala Asp Glu Gly Ser Val Leu Ala Leu Thr Val Arg Val 2930 2935 2940

Leu Asp Val Val Gin Cys Trp Leu Ala Gly Gly Gly Leu Glu Gly Thr 2945 2950 2955 2960

Lys Leu Ala Ile Val Thr Arg Gly Ala Val Pro Ala Gly Asp Gly Val 2965 2970 2975

Val His Asp Pro Ala Ala Ala Ala Val Trp Gly Leu Val Arg Ala Ala 2980 2985 2990

Gln Ala Glu Asn Pro Asp Arg Ils Val Leu Leu Asp Val Glu Pro Glu 2995 3000 3005

Als Asp Val Pro Pro Leu Leu Gly Ser Val Leu Ala Asp Gly Glu Pro 3010 3015 3020

Gln Val Ala Val Arg Gly Thr Thr Leu Ser Ile Pro Arg Leu Ala Arg 3025 3030 3035 3040

Ala Ala Arg Pro Asp Pro Ala Ala Gly Phe Lys Thr Arg Gly Pro Val 3045 3050 3055

Leu Val Thr Gly Gly Thr Gly Ser Leu Gly Gly Leu Val Ala Arg His

3250

			306	9				306	\$				307	0	
Leu	Val	Glu 307:		His	Gly	Vai	Arg 308		Leu	Val	Leu	Ala 308		Arg	Arg
Gly	Leu 309:		Als	Glu	Gly	Als 309		Asp.	Leu	Val	Thr 3100	_	Leu	Thr	Ala
leu 310:		sia	Ązp	Val	Ala 3110		Ala	ьîА	Cys	Asp 3115		Ala	Asp	Arg	As p 3120
Gln	Val	Ala	Ala	Leu 3125		Thr	Glu	Nis	Arg 3130		Ser	Ala	Val	Val 313:	
Thr	Ala	Gly	Val 314(Asp	Ala	Gly	V&l 314:		Gly	Thr	Val	Thr 315(Pro)	Asp
Arg	Leu	Ala 3155		Val	Phe	Ala	Pro 3160	_	Val	Thr	Ala	Ala 3165	•	His	Leu
Asp	Glu 317(Thr	Arg	qeA	leu 3175		Leu	Asp	Ser	Phe 3180		Val	Tyr	Ser
Ser 3185		Ser	ala	Val	Phe 3190		Gly	Ala	Gly	Ser 3198		Ser	Ţyr.	Ala	Ala 3200
Ala	Asn	Ala	Tyr	Lee 3205		Gly	leu	Met	Ala 3210		Arg	Arg	Ala	Ala 3215	
Leu	Pro	Gly	Gln 3220		Leu	Ala	Trp	Gly 3225		Trp	qaA		The 3230	Thr	Gly
Gly	Met	Ala 3235		Gly	Thr	Asp	Glu 3240		Gly	Ārģ	Ala	Arg 3245		Thr	Arg
Arg	Gly	Gly	Len	Val	Ala	Met.	Lys	Pro	Ala	Als	Gly	Leu	qzA	Leu	Phe

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Asp Ala Ile Gly Ser Gly Glu Pro Leu Leu Val Pro Ala Gln Leu 3265 3270 3275 3280

Asp Leu Arg Gly Leu Arg Ala Glu Ala Ala Gly Gly Thr Glu Val Pro 3285 3290 3295

His Leu Leu Arg Gly Leu Val Arg Ala Gly Arg Gln Gln Ala Arg Ala 3300 3305 3310

Ala Ser Thr Val Glu Glu Asn Trp Ala Gly Arg Leu Ala Gly Leu Glu 3315 3320 3325

Pro Ala Giu Arg Gly Gin Vai Leu Leu Glu Leu Vai Arg Ala Gin Val 3330 3335 3340

Ala Gly Vel Leu Gly Tyr Arg Ala Ala His Gln Vel Asp Pro Asp Gln 3345 3350 3355 3360

Gly Leu Phe Glu Ile Gly Phe Asp Ser Leu Thr Ala Ile Glu Leu Arg 3365 3370 3375

Asn Arg Leu Arg Ala Arg Thr Glu Arg Lys Ile Ser Pro Gly Val Val 3380 3385 3390

Phe Asp Ris Pro Thr Pro Ala Leu Leu Ala Ala His Leu Asn Glu Leu 3395 3400 3405

Leu Arg Lys Lys Val 3410

(2) INFORMATION FOR SEQ ID NO: 9:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 226 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single

- (D) TOPOLOGY: linear
- (ii) MULECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:

Met Ala Ile Pro Tyr Ser Ser Leu Ala Tyr Glu Leu Arg Asp Ala Val 1 5 10 15

Asn Val Val Asp Leu Asp Glu Asp Asp Val The Val Thr Ser Ile Ala
20 25 30

Glu Gly Gln Gly Gly Ala Cys Tyr His Leu Asn Arg Leu Phe His Arg 35 40 45

Leu Leu Thr Glu Leu Gly Tyr Asp Val Thr Pro Leu Ala Gly Ser Thr 50 55 60

Ala Glu Gly Arg Glu Thr Phe Gly Thr Asp Val Glu His Met Phe Asn 65 70 75 80

Leu Val Thr Leu Asp Gly Ala Asp Trp Leu Val Asp Val Gly Tyr Pro 85 90 95

Gly Pro Thr Tyr Val Glu Pro Leu Ala Val Ser Pro Ala Val Gln Thr 100 105 110

Gin Tyr Gly Ser Gin Phe Arg Leu Vel Glu Gin Glu Thr Gly Tyr Ala 115 120 125

Leu Gìn Arg Arg Gly Ala Val Thr Arg Trp Ser Val Val Tyr Thr Phe 130 135 140

The Thr Gln Pro Arg Gln Trp Ser Asp Trp Lys Glu Leu Glu Asp Asn

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145 150 155 160

Phe Arg Ala Leu Val Gly Asp Thr Thr Arg Thr Asp Thr Gln Glu Thr 165 170 175

Leu Cys Gly Arg Ala Phe Ala Asn Gly Gln Val Phe Leu Arg Gln Arg 180 185 190

Arg Tyr Leu Thr Val Glu Asn Gly Arg Glu Gln Val Arg Thr Ile Thr 195 200 205

Asp Asp Asp Glu Phe Arg Ala Leu Val Ser Arg Val Leu Ser Gly Asp 210 215 220

His Gly

Ciba-Geigy AG

CH-4002 Basel

RECEIPT IN THE CASE OF AN ORIGINAL DEPOSIT INSTERNATIONAL DEPOSITABLY AUTHORITY identified in the bosom of this page

I EDENTIFICATION OF THE DECROORGANISM provide a second states of the DESONALOR Accession number given by the INTERNATIONAL DEPOSITABLY AUTHORITY pR17-3 DSM 11114 B. SCIENTIFE: DESCRIPTION AND/OR PROPOSED TAXONOMIC DESKINATION The microorganism identified under i above was accompanied by (X) is scientific description. (X_i) * proposed taxonomic designation (Mark with a cross where applicable) HE RECEPT AND ACCEPTANCE This International Depository Assistably accepts the microorganism identified under L above, which was received by it on 1996~08~10 (Desc of the original deposit) IV. RECEIPT OF RECRIEST FOR CONVERSION The microorganism identified under I above was received by this International Depositary Authority on (date of original deposit) and a request to convent the original depose to a deposit tender the Studenest Frenty was received by it on (dasse of receipt of request for conversion) V. INTERNATIONAL DEPOSITARY AUTHORITY Name: DSMZ-DEUTSCHE SAMMLUNG VOR Signature(s) of person(s) having the power to represent the MERROCHGANISMEN UND ZEELKULTUREN GMM! international (represent Authority or of authorized official(s) Address: Maicheseder Weg 16 U. Wells D-38124 Steamschweig Dem: 1996-08-14

Form DEME-BP/4 (ante page) \$196

Where Rule 6.4 (d) applies, such due is the date on which the status of international depositary authority was acquired.

Ciba-Geigy AG

CH-4002 Basel

VIABILITY STATEMENT
issued pursuant to Role 19.2 by the
BATERNATEMAL DEPOSITARY ASSTRUMITY
identified at the bottom of this page

L DEPOSITOR		II DENTIFICATION OF THE SECROORGAMEM
	oa-Geigy AG -4002 Basel	Accession number given by the INTERNATIONAL DEPOSITABLY AUTHORSTY DSM 11114 Date of the depose of the transfer* 1996-08-10
E VEABILITY	STATEMENT	
and and a	sold microsepanism was	
***************************************	o longer viable	DESC 22-58 KNESPs*
{ } m		PERFCHMED'
EA CAMOLIKA	o longer washir IS UMDER WEICH THE VEABILITY TEST HAS BEEN	PERFORMED'
ta (2342)11894	o longer viable	PERFORMED*

Indicate the date of original deposit or, where a new deposit or a transfer has been made, the most recent relevant date (date of the new deposit or date of the transfer)

board with a cross the applicable box.

In the cures referred to in Auje 10.7(a) (ii) and (iii), well so the most recent visibility use

FM in is the information has been requested and if the results of the rest were regative.

CH-4002 Basel

RECEIPT IN THE CASE OF AN ORIGINAL DEPOSIT issued pursues to Ruic 7.1 by the INTERNATIONAL DEPOSITARY AUTHORITY identified at its bostom of this page

I. IDENTIFICATION OF THE ASCROORGANISM									
bensifesii pRi44	n sakrence given by she ICFOSITOR: - Z	Accession describe by the Intermediate althoughty DSM 11655							
II. SCIENT	iten: description and/or proposed taxonomec des	GNATION							
The microo	ganism identified under t. above was accompanied by:								
	(X) a seientific description (X) a proposed transcense desgration								
(deserbe union	s cioni mpiet abbacunie)								
us a s cen	T AND ACCEPTANCE								
This basema (Date of the	tional Depositary Authority accepts the microorganism identified t congress deposity).	index t. above, which was received by it on 1997-07-14							
IV. RECEN	T OF REQUEST FOR CONVERSION								
The micros and a reque for convers	rganism identified under 8 above was received by this lineometiconal of to convert the original deposit to a deposit under the Budapess 7 ino).	Depository Asubacity on (date of original deposit) ready was received by it on (date of receipe of request							
V. INTERN	IATIONAL DEPOSITARY AUTHORITY								
Ham:	MIKROURGAMEMEN UND ZELLKULTUREN Gmohl international Depository Authority of af substant afficial(s)								
Address	Misschenoser Weg 16 IA-38124 Braunschweig	U. Wels							
	Date: 1997-07-15								

When Safe 8.4 (d) applies, such take is the time on which the states of international depositary authority was acquired. Form DSMZ-8894 (sofe page) \$196.

CH-4002 Basel

VIABILITY STATEMENT isseed pursuant in Rule 19.2 by the INTERNATIONAL DEPOSITARY AUTHORITY identifies at the bottom of this page

i. deposit	K X	IL DENTIFICATION OF THE MICHOORGANISM
	Vovartis AG IH-4002 Basel	Accession number given by the INTERNATIONAL DEPOSITARY AUTHORITY DSM 11655 Date of the deposit of the specific 1997-07-14
ie viabi	BTV STATEMENT	
The visibility On this distr	r of the microorganism identified under N above was bested on 1. . So said microorganism was	997-07-14 1.
\$ X ()	N visite	
()	3 so kozer viable	
IV CONDA	Dions ender which the veableity test has been pe	SECHMEN
V. DYTERN	ATTOWAR DEPOSITABLY AUTHORITY	
	THE PROPERTY OF THE PROPERTY OF THE PROPERTY OF	
Name:	DSMZ-CEUTSCHE BAMMLUNG VON MBKBOORGANISMEN LIND ZELLKULTUREN GmbH	Signature(s) of persons(s) having the power in represent the International Depository Authority or of authorized official(s)
Address	Masscheroder Wag 18 D-38124 Bresmichweig	U. W. C. C. C. 1997-07-15

dedicate the same of anginal deposit or, where a new deposit or a transfer has been made, the most recent relevant date (date of the new decreas or date of the transfer).

ân time cases referred to in Rule 10.2(x) (ii) and (iii), refer to the most recem viability sest

Mark with a cross the applicable box.

Fill in 85 the information has been requested and if the results of the test were negative.

Form DSMZ-8999 (sole page) 0196

CH-4002 Basel

RECEIPT IN THE CASE OF AN ORIGINAL DEPOSIT issued pursuant to Rick 7.1 by the INTERNATIONAL DEPOSITARY AUTHORITY issueded in the bossom of this page

1. IDENTIFICATION OF THE MICROSROANISM							
ländiälission männens piese by die DEPOSITOR: DIESS	Accession number given by the INTERNATIONAL DEPOSITARY AUGMENTY DSM 11656						
II. SCIENTIFIC DESCRIPTION ANIMOR PROPOSED TAXONOMIC DESI	Cenation						
The microorganism identified under) above was accompanied by: (X) — a scientific description (X) — a proposed taxonomic designation (Mark with a cross where applicable).							
III. RECEIPT AND ACCEPTANCE							
This International Departury Authority accepts the microorganism identified a (Data of fixe original deposit).	nder Labove, which was received by it on 1997-07-14						
IV. RECEIPT (3) REQUEST FOR CONVERSION							
The microsogunism identified under I above was recrived by this international and a request in convert the original deposit to a deposit under the Sudapers I for conventant).	Depositiony Authority on (date of original deposit) restly was received by it on (date of receive of request						
V. INTERNATIONAL DEPOSITABLY AUTHORITY							
Name: DSMZ-IRVINCHE SAMPHEUNG VON MBKROORGANISMEN (MO ZELLKINTUREN OmbH Address: Muscheroder Weg 15: D-28124 Brausschweig	Signature(s) of paraoid(s) having the power to represent the International Department Authority or of authorized official(s): Compared to the						

Form (ISMZ-BP/s (axis page) 9196

³ Where Bule 6.4 (d) applies, such date is the date on which the ciasus of international depository authority was acquired

CH-4002 Basel

VIABILITY STATEMENT
issued pursuant to Ruic 16.1 by the
INTERNATIONAL DEPOSITARY AUTHORITY
identified at the bosons of this page

i derositor	B. IDENTIFICATION OF THE MICROCRGANISM
Name Novartis AG Asses: CH-4002 Basel	Accession mustifut given by the INTERNATIONAL DEPOSITABLY AUTHORITY DEM 11656 Dun of the deposit of the variety 1997-07-14
is. Viaseljty stat emen t	
The visibility of the microsorganium identified under 95 above was tested as 1.5 On that date, the soid microsorganium was (X)' visible ()' no longer visible	\$97~07 ~14 .*)
IV. CONDENONS UNDER WHICH THE VIABILITY TEST HAS BEEN FEI	EXEMPLED.
V. INTERNATIONAL DEPOSITARY AUTHORITY	
Name: DShiz-StellTSCHE SAMMLING VON MERKANGANISMEN UND ZELLKULTUREN GmbH Address: Mantherodes Weg 15 D-38124 Braunschweig	Signature(s) of person(s) having the power to represent the international Depository Authority or of aethorized official(s). U. ULC. Date: 1987-07-15

indicate the data of original disposit or, where a new deposit or a bazzeler has been made, the most extent reservant date (date of the serv deposit or date of the bazzeler).

in the cases referred to in Role 10 2(a) (ii) and (iii), refer to the most recent vishilly test.

Mark with a cross the applicable bux.

Form DSNIZ-ISP/9 (sede page) 0196

Pill in if the information has been requested and if the results of the test were negative.

CH-4002 Basel

RECEIPT IN THE CASE OF AN ORIGINAL DEPOSIT issued pursuant to Ruin 7.1 by the INTERNATIONAL DEPOSITARY AUTHORITY international of this page

I. IDENTIFICATION OF THE ARCROGROAMESM				
lormsfessio pNE112	n reference given by the DEPOSITER:	Accession success process by the INTERNATIONAL DEPOSITARY AUTHORITY: DSM 11657		
IL SCIENTIFIC DESCRIPTION AND/OR PROPOSED TAXONOMIC DESIGNATION				
The microstypenen identified under 1 share was accompanied by: (X) a seinesific description (X) a proposed execution designation (Mark with a cross where explicable).				
in recept and acceptance				
This international Depositary Authority accepts the microsografism identified under 1. above, which was received by it on 1997~07-14 (Date of the original deposity)				
IV. RECEIPT OF REQUEST FOR CONVERSION				
The microsegueism identified under (above was received by this International Depository Assimption on (date of exiginal deposit) and a request to convert the original deposit to a deposit under the Budapest Treaty was received by 8 on (date of receipt of request for convention).				
V. INTERNATIONAL DEPOSITARY AUTISORITY				
Piane: Address:	DSMZ-DEUTSCHE SAMMLUNG VON MIKROORGANISMEN UND ZELLKULTUREN GmbH Maschender Weg ib D-38124 Brautschweig	Signature(s) of persons) basing the power to represent the international Depository Authority of oil authopited (officialis). U. U.Z.Z.		
		Ome: 1997-07-15		

Form DSMZ-BP6 (2002 page) 0196

When Note 6.4 (d) applies, such dass is the date on which the status of international depositiony authority was acquired.

CH-4002 Basel

VIABILITY STATEMENT
issued pursues to Rule 10.2 by the
INTERNATIONAL DEPOSITARY AUTHORITY
identified at the bossom of this page

). DEPOSITIOR		II. IDENTIFICATION OF THE MICROSIGANISM	
Nems: Address:	Novartis AG CH-4002 Basel	Accession member given by the ONTERNATIONAL DEPOSITABLY AUTHORITY DSN 11657 Date of the deposit of the transfer 1937-07-14	
i Viabi	LITY STATEMENT		
ihe viskili In that ¢a	ity of the microorpanism identified under 11 above was usued on 1 te, 8% said microorpanism was	997~07~143	
()	Sign with the sign of the sign		
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Y. COND	ficans under which the viability test has been be	XFORMED.	
/. Intero	VATIONAL DEPOSITABLY AUTHORITY		
V. INTERN	YATIONAL DEPOSITARY AUTHORITY DEMO-DEUTRORE SAMORLUNG VON MIKROORGANISMEN UND ZELLKELTEREN GM8H	Signature(s) of person(s) having the power to represent the last management of persons Authority or of substricts official(s).	
 w	DSM2-DESTRONE SANGRUNG VON	Signature(s) of person(s) having the power to represent the learnaminal Depository Authority or of enthusized official(s).	

Indicese the date of original deposit or, where a new deposit or a transfer has been made, the most recent resevant tists (date of the new deposit or date of the namely).

Mark with a cross the applicable box.

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In the cases referred to in Rule 19.2(a) (ii) and (iii), refer to the most recent richility seat

Fill in if the information has been requested and if the results of the test were negative.

What is claimed is:

- 1. A DNA fragment from the genome of Amycolatopsis mediterranei which comprises a DNA region which is involved directly or indirectly in the gene cluster responsible for rifamycin synthesis, including the adjacent DNA regions to the right and left which, by reason of their function in connection with rifamycin biosynthesis, qualify as constituent of this rifamycin gene cluster; and functional fragments, derivatives or constituents thereof.
- A DNA fragment according to claim 1, which is directly or indirectly involved in the gene cluster responsible for rifamycin synthesis.
- A DNA fragment according to claim 1, which comprises sequence portions which code for a polyketide synthase or an enzymatically active domain thereof.
- A DNA fragment according to claim 1, which comprises SEQ ID NO 1 or SEQ ID NO
 3 or at least 15 consecutive nucleotides therefrom.
- 5. A DNA fragment according to claim 1, wherein said fragment comprises one or more of the partial nucleotide sequences depicted in SEQ ID NOS 1 and/or 3, or functional fragments thereof, and all other DNA sequences in the vicinity of this sequence which can, by reason of homologies which are present, be regarded as structural or functional equivalents and are therefore able to hybridize with this sequence.
- 6. A DNA fragment according to claim 1, wherein said fragment comprises a nucleotide sequence selected from the group consisting of ORF A, B, C, D, E and F or functional fragments thereof, or encodes one or more of the proteins or polypeptides, or functional derivatives thereof, depicted in SEQ ID NOS 4 to 9.
- 7. A method for identifying, isolating and cloning a DNA fragment according to claim 1.

- 8. A method according to claim 7, which comprises the following steps:
 - setting up of a genomic gene bank,
 - screening of this gene bank with the assistance of the DNA sequences according to the invention, and
 - isolation of the clones identified as positive.
- 9. The use of a DNA fragment according to claim 1 in the production of ansamycins or precursors thereof; including those in which the aliphatic bridge is connected only at one end to the aromatic nucleus.
- 10. The use of a DNA fragment according to claim 1 in the production of rifamycin, rifamycin analogues or precursors thereof.
- 11. The use of a DNA fragment according to claim 1 for inactivating or modifying genes of ansamycin biosynthesis.
- 12. The use of a DNA fragment according to claim 1 for inactivating or modifying genes of rifamycin biosynthesis, or the biosynthesis of rifamycin analogues.
- 13. The use of a DNA fragment according to claim 1 for constructing mutated actinomycetes strains from which the natural rifamycin or ansamycin biosynthesis gene cluster in the chromosome has been partly or completely deleted.
- 14. The use of DNA fragments according to claim 1 for assembling a library of polyketide synthases.
- 15. The use of the polyketide synthases according to claim 14 for assembling a library of polyketides.
- 16. A polyketide synthase from *Amycolatopsis mediterranei* which is directly or indirectly involved in rifamycin synthesis; and functional constituents or domains thereof.

- 17. The use of the polyketide synthase according to claim 16 for synthesizing ansamycins.
- 18. The use of polyketide synthases according to claim 14 for synthesizing a library of ansamycins.
- 19. A hybrid vector comprising a DNA fragment according to claim 1.
- 20. A hybrid vector comprising an expression vector comprising a DNA fragment according to claim 1.
- A host organism comprising a hybrid vector according to claim 19.
- 22. A hybridization probe comprising a DNA fragment according to claim 1.
- 23. The use of the hybridization probe according to claim 22 for identifying DNA fragments involved in the biosynthesis of ansamycins.

INTERNATIONAL SEARCH REPORT

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